

10/661458

INVENTOR SEARCH

>> fil cap1; d que 11; d que 145
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FILE COVERS 1907 - 14 Dec 2006 VOL 145 ISS 25
FILE LAST UPDATED: 13 Dec 2006 (20061213/ED)

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"OBI" IS DEFAULT SEARCH FIELD FOR 'CAPLUS' FILE

L1 1 SEA FILE=CAPLUS ABB=ON US2003-661458/APPS

L2 141 SEA FILE=CAPLUS ABB=ON PACE G7/AU
L3 11003 SEA FILE=CAPLUS ABB=ON SMITH M7/AU
L5 1 SEA FILE=REGISTRY ABB=ON MORPHINE/CN
L6 1 SEA FILE=REGISTRY ABB=ON FENTANYL/CN
L7 1 SEA FILE=REGISTRY ABB=ON SUFENTANIL/CN
L8 1 SEA FILE=REGISTRY ABB=ON ALFENTANIL/CN
L9 1 SEA FILE=REGISTRY ABB=ON OXYMORPHONE/CN
L10 1 SEA FILE=REGISTRY ABB=ON HYDROMORPHONE/CN
L11 1 SEA FILE=REGISTRY ABB=ON OXYCODONE/CN
L12 31087 SEA FILE=CAPLUS ABB=ON (L5 OR L6 OR L7 OR L8 OR L9 OR L10)
L13 1073 SEA FILE=CAPLUS ABB=ON L11
L14 12914 SEA FILE=CAPLUS ABB=ON OPIOIDS/CT
L15 1209 SEA FILE=CAPLUS ABB=ON L14(L)KAPPA/OBI
L16 1944 SEA FILE=CAPLUS ABB=ON L14(L)MU/OBI
L17 56591 SEA FILE=CAPLUS ABB=ON AGONISTS/OBI
L18 368 SEA FILE=CAPLUS ABB=ON L15(L)L17
L19 454 SEA FILE=CAPLUS ABB=ON L16(L)L17
L20 39125 SEA FILE=CAPLUS ABB=ON DRUG INTERACTIONS-OLD,NT/CT
L21 4450 SEA FILE=CAPLUS ABB=ON DRUG DELIVERY SYSTEMS-OLD,CT(L)COMB?/OB
L22 1
L23 16989 SEA FILE=CAPLUS ABB=ON COMBINATION CHEMOTHERAPY/CT
L24 5480 SEA FILE=CAPLUS ABB=ON COMB?/OBI(L)PHARMAC?/OBI
L25 552 SEA FILE=CAPLUS ABB=ON (L12 OR L19)(L)(COMB?/OBI OR COADMIN?/O
BI OR CODRUG?/OBI OR CONCOMITANT?/OBI OR CONCURRENT?/OBI OR
BLEND?/OBI OR MIXTURES?/OBI)
L26 82 SEA FILE=CAPLUS ABB=ON (L13 OR L18)(L)(COMB?/OBI OR COADMIN?/O
BI OR CODRUG?/OBI OR CONCOMITANT?/OBI OR CONCURRENT?/OBI OR

10/661458

L45 5 SEA FILE=CAPLUS ABB=ON ((L42 AND L43) OR ((L12 OR L19) AND
(L13 OR L18) AND (L37 OR L38 OR L39 OR L40))) AND (L2 OR L3)

>> s 11.145

L210 5 (L1 OR L45)

>> fil embase; d que 181

FILE 'EMBASE' ENTERED AT 11:10:14 ON 14 DEC 2006
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FILE COVERS 1974 TO 13 Dec 2006 (20061213/ED)

EMBASE is now updated daily. SDI frequency remains weekly (default)
and biweekly.

This file contains CAS Registry Numbers for easy and accurate
substance identification.

L46 83 SEA FILE=EMBASE ABB=ON PACE G7/AU
L47 8120 SEA FILE=EMBASE ABB=ON SMITH M7/AU
L48 53452 SEA FILE=EMBASE ABB=ON MORPHINE/CT
L49 26736 SEA FILE=EMBASE ABB=ON FENTANYL/CT OR FENTANYL CITRATE/CT
L50 4395 SEA FILE=EMBASE ABB=ON SUFENTANIL/CT OR SUFENTANIL CITRATE/CT
L51 4482 SEA FILE=EMBASE ABB=ON ALFENTANIL/CT
L52 805 SEA FILE=EMBASE ABB=ON OXYMORPHONE/CT
L53 2957 SEA FILE=EMBASE ABB=ON HYDROMORPHONE/CT
L54 3754 SEA FILE=EMBASE ABB=ON OXYCODONE/CT
L72 493 SEA FILE=EMBASE ABB=ON L54(L)(CB OR IT)/CT
L80 10397 SEA FILE=EMBASE ABB=ON (L48 OR L49 OR L50 OR L51 OR L52 OR
L53)(L)(CB OR IT)/CT
L81 5 SEA FILE=EMBASE ABB=ON (L46 AND L47) OR (L80 AND L72 AND
L47))

>> fil drugu; d que 196

FILE 'DRUGU' ENTERED AT 11:10:15 ON 14 DEC 2006

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FILE LAST UPDATED: 11 DEC 2006 <20061211/UP>

>>> DERMENT DRUG FILE (SUBSCRIBER) <<<

>>> FILE COVERS 1983 TO DATE <<<

>>> THESAURUS AVAILABLE IN /CT <<<

L5 1 SEA FILE=REGISTRY ABB=ON MORPHINE/CN
L6 1 SEA FILE=REGISTRY ABB=ON FENTANYL/CN
L7 1 SEA FILE=REGISTRY ABB=ON SUFENTANIL/CN
L8 1 SEA FILE=REGISTRY ABB=ON ALFENTANIL/CN
L9 1 SEA FILE=REGISTRY ABB=ON OXYMORPHONE/CN
L10 1 SEA FILE=REGISTRY ABB=ON HYDROMORPHONE/CN
L11 1 SEA FILE=REGISTRY ABB=ON OXYCODONE/CN

1

2

10/661458

L111 3147 SEA FILE=MPIX ABB=ON MORPHINE/B1,ABEX OR FENTANYL/B1,ABEX OR
ALFENTANIL/B1,ABEX OR SUFENTANIL/B1,ABEX OR OXYMORPHONE/B1,ABEX
OR MRZ2593/B1,ABEX OR MRZ 2593/B1,ABEX OR HYDROMORPHONE/B1,ABEX

X
L112 4 SEA FILE=MPIX ABB=ON OXYCODONE/CT
L113 431 SEA FILE=MPIX ABB=ON L112/DCN
L114 4 SEA FILE=MPIX ABB=ON (RA800/SDCN OR RACDH7/SDCN OR RA0FC0/SDC
N OR R06854/SDCN OR R16303/SDCN OR 103043-1-2-0/DCSE OR 758270-1-0-0/DCSE)
103043-1-1-0/DCSE OR 103043-1-2-0/DCSE OR 758270-1-0-0/DCSE)
L115 435 SEA FILE=MPIX ABB=ON L114 OR L113
L116 513 SEA FILE=MPIX ABB=ON OXYCODONE/B1,ABEX
L117 198 SEA FILE=MPIX ABB=ON OPIOID/B1,ABEX
L118 186 SEA FILE=MPIX ABB=ON KAPPA/B1,ABEX (L1M) OPIOID/B1,ABEX
L119 12146 SEA FILE=MPIX ABB=ON B14-L01/MC OR C14-L01/MC
L120 100 SEA FILE=MPIX ABB=ON L117(2A)AGONIST#/B1,ABEX OR (L117 AND
L119)
L121 102 SEA FILE=MPIX ABB=ON L118(2A)AGONIST#/B1,ABEX OR (L118 AND
L119)
L122 486502 SEA FILE=MPIX ABB=ON (M782 OR P667)/M0,M1,M2,M3,M4,M5,M6 OR
A61K045/IPC OR (B12-C09 OR C12-C09 OR B14-S09 OR C14-S09)/MC
L123 4 SEA FILE=MPIX ABB=ON (L108 OR L109) AND (L111 OR L120) AND
(L115 OR L116 OR L121) AND L122

>> s 1110.1123

L211 4 (L110 OR L123)

>> fil medi; d que 1163

FILE 'MEDLINE' ENTERED AT 11:10:19 ON 14 DEC 2006

FILE LAST UPDATED: 13 Dec 2006 (20061213/UP), FILE COVERS 1950 TO DATE.

In preparation for the annual MEDLINE reload, the National Library of
Medicine (NLM) has suspended delivery of regular updates as of November
15, 2006. In-process and in-data-review records will resume delivery
on November 21, 2006, and will continue to be added to MEDLINE until
December 17, 2006.

On December 17, 2006, all regular MEDLINE updates from November 15 to
December 16 will be added to MEDLINE, along with 2007 Medical Subject
Headings (MeSH(R)) and 2007 tree numbers.

The annual reload will be available in early 2007.

This file contains CAS Registry Numbers for easy and accurate
substance identification.

L144 94 SEA FILE=MEDLINE ABB=ON PACE G7/AU
L145 10732 SEA FILE=MEDLINE ABB=ON SMITH M7/AU
L146 0 SEA FILE=MEDLINE ABB=ON L144 AND L145
L147 28104 SEA FILE=MEDLINE ABB=ON MORPHINE/CT
L148 10382 SEA FILE=MEDLINE ABB=ON FENTANYL-NT/CT
L149 294 SEA FILE=MEDLINE ABB=ON OXYMORPHONE/CT
L150 704 SEA FILE=MEDLINE ABB=ON HYDROMORPHONE/CT
L151 540 SEA FILE=MEDLINE ABB=ON OXYCODONE/CT
L152 124891 SEA FILE=MEDLINE ABB=ON LUNG DISEASES, OBSTRUCTIVE-NT/CT
L153 5936 SEA FILE=MEDLINE ABB=ON BRONCHIECTASIS-NT/CT

3

4

L154(57086)SEA FILE+MEDLINE ABB+ON TUBERCULOSIS, PULMONARY+NT/CT
 L155(34601)SEA FILE+MEDLINE ABB+ON BRONCHOPNEUMONIA/CT
 L156(3610)SEA FILE+MEDLINE ABB+ON LARYNGITIS-NT/CT
 L157(11628)SEA FILE+MEDLINE ABB+ON SINUSITIS-NT/CT
 L158(13172)SEA FILE+MEDLINE ABB+ON PULMONARY FIBROSIS/CT
 L159(1561)SEA FILE+MEDLINE ABB+ON SARCOIDOSIS, PULMONARY/CT
 L160(113814)SEA FILE+MEDLINE ABB+ON LUNG NEOPLASMS-NT/CT
 L161(12706)SEA FILE+MEDLINE ABB+ON SLEEP APNEA SYNDROME-NT/CT
 L162(0)SEA FILE+MEDLINE ABB+ON ((L144 OR L145) AND (L147 OR L148 OR
 L149 OR L150 OR L151) AND (L152 OR L153 OR L154 OR L155 OR
 L156 OR L157 OR L158 OR L159 OR L160 OR L161)
 L163(0)SEA FILE+MEDLINE ABB+ON L146 OR L162

> dup rem 196,1210,1211,181
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PROCESSING COMPLETED FOR L96
 PROCESSING COMPLETED FOR L210
 PROCESSING COMPLETED FOR L211
 PROCESSING COMPLETED FOR L81

L212 16 DUE REM L96 L210 L211 L81 (7 DUPLICATES REMOVED)
 ANSWERS '1-9' FROM FILE DRUGU
 ANSWERS '10-13' FROM FILE CAPLUS
 ANSWER '14' FROM FILE WPIX
 ANSWERS '15-16' FROM FILE EMBASE

> d ibib ed abs 1-16

L212 ANSWER 1 OF 16 DRUGU COPYRIGHT 2006 THE THOMSON CORP on STN DUPLICATE 2
 ACCESSION NUMBER: 2005-27268 DRUGU P S Full-text
 TITLE: Ventilatory responses of healthy subjects to intravenous
 combinations of morphine and oxycodone under imposed
 hypercapnic and hypoxicemic conditions.
 AUTHOR: Ladd L A; Kam P C; Williams D B; Wright A W E; Smith M
 T; Mather L E
 CORPORATE SOURCE: Univ.Sydney; Sigma-Pharmaceuticals; Univ.Queensland
 LOCATION: Brisbane; Melbourne, Austr.
 SOURCE: Br.J.Clin.Pharmacol. (59, No. 5, 524-35, 2005) 5 Fig. 2 Tab.
 37 Ref.
 CODEN: BCPHBM ISSN: 0306-5251
 AVAIL. OF DOC.: Department of Anaesthesia and Pain Management, University of
 Sydney at Royal North Shore Hospital, St Leonards, NSW 2065,
 Australia. (L.E.M.). (e-mail: lmather@med.usyd.edu.au).
 LANGUAGE: English
 DOCUMENT TYPE: Journal
 FIELD AVAIL.: AB; LA; CT
 FILE SEGMENT: Literature
 AN 2005-27268 DRUGU P S Full-text

AB I.v. infusions of morphine sulfate (NOR) or oxycodone HCl (OXY) or their combination decreased the hypercapnic response and VES5 (i.e., mean minute ventilation at $\text{PTCO}_2=55 \text{ mmHg}$) to a similar degree in a randomized, placebo-controlled, double-blind, crossover study of 12 male volunteers. There was no consistent treatment effect on the hypoxicemic response. NOR was associated with drowsiness, tingling, warm feeling, itching, and nausea. These findings suggest that no unexpected or disproportionate effects are expected of NOR and OXY treatments that might impede their use in combination for pain management.

ABEX Methods 12 Male volunteers (aged 18-45 yr) randomly received 1-hr i.v. infusions of placebo, NOR (5, 10, and 15 mg, M15, M10, and M5, respectively), OXY (5, 7.5, 10, and 15 mg, Q5-015, respectively), or their combination in dose ratios of 1:2, 1:1, and 2:1, in a crossover manner. Results Subjective side-effects increased with increasing OXY doses. Drowsiness, tingling, and warm feeling were mostly mild and random, although some subjects tended to experience recurring side-effects (e.g., itching or nausea). A consistent treatment effect was not demonstrated for slope or intercept of the hypoxicemic response. There was a consistent and similar decrease in slope of the hypercapnic response during all active drug treatments (DT), with general recovery after treatment. There was also a consistent decrease of VES5 during all treatments, with partial recovery after DT, but not between active DT. During DT, VES5 decreased to a mean of 74% of the respective values before DT (74%, 65%, 69%, 68%, and 61% for M15, M10/05, M7.5/07.5, M5/10, and Q5, respectively). After DT, mean values of VES5 were 75%, 73%, 78%, 76%, and 75% of the respective values before DT. Drug and metabolite AUC 0-120 hr were linearly proportional to dose and did not differ between drugs. Although there were differences in mean plasma drug concentrations between subjects, there were no differences between treatments during infusion; differences were found between treatments after infusion, with concentrations being directly correlated with the OXY dose. VES5 was the most sensitive ventilatory response variable for comparing the individuals and treatments in relation to drug plasma concentrations. (ABD/Y230)

L212 ANSWER 2 OF 16 DRUGU COPYRIGHT 2006 THE THOMSON CORP on STN DUPLICATES 4
 ACCESSION NUMBER: 2000-12785 DRUGU P S Full-text
 TITLE: Co-administration of sub-antinociceptive doses of oxycodone and morphine produces marked antinociceptive synergy with reduced CNS side effects in rats.
 AUTHOR: Ross F B; Wallis S C; Smith M T
 CORPORATE SOURCE: Univ.Queensland
 LOCATION: Brisbane, Austr.
 SOURCE: Pain (84, No. 2-3, 421-28, 2000) 4 Fig. 24 Ref.
 CODEN: PAINDB ISSN: 0374-3959
 AVAIL. OF DOC.: School of Pharmacy, The University of Queensland, St Lucia, Brisbane, Queensland 4072, Australia. (M.T.S.). (e-mail: Mares.smith@pharmacy.uq.edu.au).

LANGUAGE: English
 DOCUMENT TYPE: Journal
 FIELD AVAIL.: AB; LA; CT
 FILE SEGMENT: Literature
 AN 2000-12785 DRUGU P S Full-text
 AB The effects of i.c.v., i.p. and s.c. oxycodone hydrochloride (Boots) and morphine hydrochloride on nociception were studied in rats. Co-administration of oxycodone and morphine increased the levels of antinociception. Behaviorally rats that received equi-potent doses of either opioid alone were markedly sedated. The results suggested that co-administration of sub-analgesic doses of oxycodone and morphine could provide excellent pain relief with a reduction in opioid related CNS side effects.

ABEX Marked antinociceptive synergy was seen in Sprague-Dawley and Dark Agouti rats following sub-antinociceptive doses of oxycodone or morphine, irrespective of whether they were given i.c.v., i.p. or s.c. Sub-antinociceptive doses of either oxycodone or morphine alone to rats produced levels of antinociception similar to pre-dosing baseline levels. In Sprague-Dawley rats i.c.v. oxycodone at 40 nmol and morphine at 15 nmol caused a rapid onset (by 10 min) of maximum possible antinociception (MPE) which decreased relatively slowly (mean level of antinociception greater than 50% of MPE at 3 hr). Pretreatment with naloxomazine or nor-BNI 24 hr prior to i.c.v. oxycodone 40 nmol plus morphine 15 nmol resulted in a decrease in the levels of antinociception. In Dark Agouti rats i.p. oxycodone at 571 nmol plus morphine at 621 nmol resulted in 100% MPE by 10 min. The mean levels of antinociception remained high for the 1st 2 hr of the experimental period and then decreased to 65% MPE by 3 hr postdosing. Oxycodone 571 nmol i.p. with 310 nmol morphine or oxycodone 285 nmol plus 621 nmol morphine resulted in maximum antinociception by 15 min but the duration of action was reduced to 2 hr. Co-administration of oxycodone or morphine in sub-antinociceptive doses neither strain of rat showed any adverse behavioral effects such as sedation, incontinence or catalepsy. In Dark Agouti rats the ED50 doses of s.c. oxycodone and morphine were 2.8 and 8.5 mg/kg, respectively. Behaviorally rats given single s.c. morphine or oxycodone in doses larger than the ED50 were sedated. Co-administration of sub-antinociceptive doses of oxycodone and morphine produced synergistic levels of pain relief. (LB)

L212 ANSWER 3 OF 16 DRUGU COPYRIGHT 2006 THE THOMSON CORP on STN DUPLICATE 6
 ACCESSION NUMBER: 1994-22863 DRUGU P S Full-text
 TITLE: The antinociceptive potencies of oxycodone, noroxycodone and morphine after intrathecerebroventricular administration to rats.
 AUTHOR: Lao K P; Smith M T
 CORPORATE SOURCE: Univ.Queensland
 LOCATION: Brisbane, Australia
 SOURCE: Life Sci. (54, No. 17, 1229-36, 1994) 2 Fig. 1 Tab. 20 Ref.
 CODEN: LIFSAK ISSN: 0024-3205
 AVAIL. OF DOC.: Department of Pharmacy, The University of Queensland, St Lucia, Queensland 4072, Australia. (M.T.S.).

LANGUAGE: English
 DOCUMENT TYPE: Journal
 FIELD AVAIL.: AB; LA; CT
 FILE SEGMENT: Literature
 AN 1994-22863 DRUGU P S Full-text
 AB In rats, i.c.v. administration of noroxycodone (NOR, Du-Pont-Merck) or oxycodone HCl (OXY, Sigma-Chemical) had a more potent antinociceptive effect than that of i.c.v. morphine HCl (MOR). Administration of i.c.v. naloxone HCl (Sigma-Chemical) abolished the antinociceptive response produced by the subsequent administration of OXY, NOR or MOR, indicating that the antinociceptive effects of these 3 drugs are mediated by opioid receptors. NOR also produced excitatory effects throughout the antinociceptive range, the severity of which was reduced, but not abolished, by prior administration of i.c.v. naloxone. As excitatory effects have not been observed in patients receiving OXY, it is unlikely that NOR contributes to the analgesic activity of OXY administered systemically.

ABEX In male Sprague-Dawley rats (250 g), the ED50 value for i.c.v. NOR was 34 nmol. Corresponding ED50 values for i.c.v. OXY and NOR were 78 and 200 nmol, respectively. Antinociceptive potencies of OXY and NOR relative to MOR, estimated using the ED50 values, were 0.44 and 0.17, respectively. After i.c.v. MOR, the antinociceptive response comprised 2 distinct phases. During phase 1, antinociception commenced at 15-30 min,

peaked at 45-60 min and decreased at 75 min. Phase 2 antinociception peaked at 90 min and decreased throughout the remainder of the 3-hr observation period. During phase 2 antinociception, rats were incontinent. Only phase 1 antinociception was observed in rats given OXY. Onset of antinociception was very rapid with peak values occurring at 7-15 min post-dosing. When NOR was administered, 2 antinociceptive phases were observed in a manner analogous to that observed after i.c.v. MOR. Time to achieve maximum antinociception was significantly shorter for OXY (9.3 min) than for MOR (31.8 min) or NOR (34.6 min). At equieffective doses, the mean duration of antinociception was significantly shorter for i.c.v. OXY (114 min) than for i.c.v. MOR and NOR (160 min). Naloxone (55 nmol) given 15 min prior to i.c.v. opioid agonist significantly reduced the antinociceptive response of the respective opioid agonist administered alone. NOR also produced allodynia, excessive facial grooming, tremor, Straub tail and myoclonic jerks. The severity of these effects was reduced but not eliminated by subsequent naloxone. Grand mal seizures then death occurred in 2 rats given 432 nmol of NOR. (SAB)

L212 ANSWER 4 OF 16 DRUGU COPYRIGHT 2006 THE THOMSON CORP on STN
 ACCESSION NUMBER: 2000-44276 DRUGU P Full-text
 TITLE: Incomplete, asymmetric, and route-dependent cross-tolerance between oxycodone and morphine in the Dark Agouti rat.
 AUTHOR: Nielsen C K; Ross F B; Smith M T
 CORPORATE SOURCE: Univ.Queensland
 LOCATION: Brisbane, Austr.
 SOURCE: J.Pharmacol.Exp.Ther. (295, No. 1, 91-99, 2000) 4 Fig. 4 Tab.
 29 Ref.
 CODEN: JPETAB ISSN: 0022-3565
 AVAIL. OF DOC.: School of Pharmacy, The University of Queensland, St. Lucia, Queensland 4072, Australia. (M.T.S.). (e-mail: m.smith@pharmacy.uq.edu.au).

LANGUAGE: English
 DOCUMENT TYPE: Journal
 FIELD AVAIL.: AB; LA; CT
 FILE SEGMENT: Literature
 AN 2000-44276 DRUGU P Full-text
 AB The antinociceptive effects of bolus i.v. or i.c.v. oxycodone HCl (OX, Tasmanian Alkaloids) or morphine sulfate (MP) were determined OX- and MP-tolerant rats. In MP-tolerant rats, i.c.v. OX did not induce cross-tolerance whereas i.v. OX induced a low degree of cross-tolerance. In OX-tolerant rats, both i.c.v. and i.v. induced a high degree of cross-tolerance. It was concluded that after parenteral but not supraspinal administration, OX is metabolized to a mu-opioid agonist metabolite, thereby explaining asymmetric and incomplete cross-tolerance between OX and MP.

ABEX Methods Dark Agouti rats (200 g) received i.v. infusion of OX (2.5 or 5 mg/day) or MP (10 or 20 mg/day) until rats were completely tolerant followed by 12-hr washout period. OX-tolerant, MP-tolerant and drug-naive rats received either bolus i.v. OX (79-1585 nmol) or MP (350-3504 nmol) or bolus i.c.v. OX (22-132 nmol) or MP (18-150 nmol). Results Complete antinociceptive tolerance was produced by 48 hr in naive rats following chronic i.v. infusion of OX (2.5 mg/day) and MP (10 mg/day). Chronic i.v. infusion of OX (5 mg/day) induced tolerance in naive, OX-tolerant and OX-tolerant rats after 72 hr, 48 hr and 8 hr, respectively. Chronic i.v. infusion of MP (20 mg/day) induced tolerance in naive, OX-tolerant and MP-tolerant rats after 84 hr, 36 hr and 12 hr, respectively. Equipotent antinociception was produced by chronic i.v. OX and MP in doses of 2.5 mg/day and 10 mg/day, respectively, and tolerance was established over a similar time frame. In MP-tolerant rats, i.c.v. OX did not affect the dose-response curve or ED50 of i.c.v. OX, whereas

i.c.v. MP increased the ED50 of i.c.v. MP. In OX-tolerant and MP-tolerant rats, i.c.v. MP caused a rightward shift in the dose-response curve of i.c.v. MP and increased the ED50 of i.c.v. MP by 1.9-fold and 2.6-fold, respectively. Rats that received i.c.v. or i.v. MP or OX were sedated, whereas rats that received i.c.v. MP experienced urinary incontinence. In MP-tolerant rats, i.v. OX and i.v. MP increased the ED50 of OX. Similarly in OX-tolerant rats, i.v. MP and i.v. OX increased the ED50 of OX. I.v. OX produced a lower degree of tolerance in MP-tolerant rats than did i.v. MP in OX-tolerant rats (33.7% vs. 71.3%). (NKK)

L212 ANSWER 5 OF 16 DRUGU COPYRIGHT 2006 THE THOMSON CORP on STN
ACCESSION NUMBER: 1998-04921 DRUGU P Full-text

TITLE: The intrinsic antinociceptive effects of oxycodone appear to be K-opioid receptor mediated.
AUTHOR: Ross F B; Smith M T
CORPORATE SOURCE: Univ.Queensland
LOCATION: Brisbane, Austr.
SOURCE: Pain (73, No. 2, 151-57, 1997) 5 Fig. 33 Ref.
CODEN: PAINDB ISBN: 0304-3959
AVAIL. OF DOC.: School of Pharmacy, Steele Building, The University of Queensland, St. Lucia, Brisbane, Queensland 4072, Australia.
(E-mail: mareas.smith@pharmacy.uq.edu.au).

LANGUAGE: English
DOCUMENT TYPE: Journal
FIELD AVAIL.: AB; LA; CT
FILE SEGMENT: Literature
AN 1998-04921 DRUGU P Full-text

AB The Authors' previous studies in the Sprague-Dawley rat showed that the intrinsic antinociceptive effects of oxycodone are naloxone reversible in a manner analogous to morphine but, in contrast to morphine, oxycodone's antinociceptive effects have a rapid onset of maximum effect, comprise 1 (not 2) antinociceptive phases and are of relatively short duration. This study used a range of selective opioid receptor antagonists to identify the major class of opioid receptors mediating the intrinsic antinociceptive effects of oxycodone following its i.c.v. administration to rats. The data strongly suggested that the antinociceptive actions of oxycodone are mediated by kappa-opioid receptors, in contrast to morphine which interacts primarily with u-opioid receptors.

ABEX A range of selective opioid receptor antagonists were given to adult male Sprague Dawley rats (200 +/- 20 g). The intrinsic antinociceptive effects of oxycodone (171 nmol) were not attenuated by i.c.v. administration of (i) naloxazine (1 nmol), a u1-selective opioid receptor antagonist, or (ii) naltrindole (2.2 nmol), a delta-selective opioid receptor antagonist, in doses that completely attenuated the intrinsic antinociceptive effects of equipotent doses of the respective u1 and delta-opioid agonists, morphine (78 nmol) and enkephalin-2-D-Pen-5-Pen (DPDPE, 55 nmol). Although beta-funaltrexamine (B-FNA, 4 nmol) attenuated the antinociceptive effects of oxycodone (171 nmol i.c.v.), it also attenuated the antinociceptive effects of morphine (78 nmol) and bremazocine (57 nmol; kappa-opioid agonist) indicative of non-selective antagonism. Importantly, the antinociceptive effects of oxycodone (171 nmol i.c.v.) were markedly attenuated by the prior i.c.v. administration of the selective kappa-opioid receptor antagonist, norbinaltorphimine, in a dose (0.3 nmol) that did not attenuate the antinociceptive effects of an equipotent dose of i.c.v. morphine (78 nmol). (PMH)

L212 ANSWER 6 OF 16 DRUGU COPYRIGHT 2006 THE THOMSON CORP on STN
ACCESSION NUMBER: 1994-27116 DRUGU P Full-text

TITLE: Serum protein binding of oxycodone and morphine.

AUTHOR: Wright A W E; Leow K P; Cromond T; Smith M T

CORPORATE SOURCE: Univ.Queensland

LOCATION: Brisbane, Australia

SOURCE: Aust.J.Hosp.Pharm. (24, No. 2, 206, 1994)

CODEN: AUHPA1 ISSN: 0310-6810

AVAIL. OF DOC.: Dept. of Pharmacy, The University of Queensland, Brisbane, Queensland, 4072, Australia.

LANGUAGE: English

DOCUMENT TYPE: Journal

FIELD AVAIL.: AB; LA; CT

FILE SEGMENT: Literature

AN 1994-27116 DRUGU P Full-text

AB The study aim was to determine the extent of serum protein binding of oxycodone (OX) and morphine (MO) in HSA and human alpha1-acid glycoprotein (AAG). OX and MO bound primarily to HSA, although both drugs bound to AAG with a higher affinity than to albumin. A decrease in temperature or an increase in pH significantly increased the protein binding of both OX and MO. The serum protein binding of both opioids was independent of drug concentration. It is unlikely that changes in serum protein concentrations associated with disease states such as renal or hepatic failure would alter the pharmacological effects of OX or MO due to the normally low extent of binding of both drugs. (congress abstract).

ABEX Methods Serum protein binding was determined in-vitro by ultrafiltration. Binding studies were also performed using both purified HSA and AAG. Results OX and MO bound primarily to albumin although both drugs bound to AAG with a higher affinity than to albumin. At physiological pH and temperature, the mean serum protein binding of OX and MO were 45.1% and 35.3%, respectively. A decrease in temperature (from 37 deg to 23 deg) or an increase in pH (from 7.4 to 7.75-7.85) significantly increased the protein binding of both OX and MO, underlining the necessity to conduct protein binding studies at physiological pH and temperature. The serum protein binding of both opioids was independent of drug concentration in the therapeutic range (5-100 ng/ml), but was dependent on the protein concentration. In serum containing albumin and AAG concentrations within the normal ranges, the binding of OX to albumin and AAG would be in the ranges 31-39% and 5-10%, respectively, and the binding of morphine would be 26-34% and 4-5%, respectively. (SAB)

L212 ANSWER 7 OF 16 DRUGU COPYRIGHT 2006 THE THOMSON CORP on STN

ACCESSION NUMBER: 1993-52909 DRUGU P Full-text

TITLE: Determination of the Serum Protein Binding of Oxycodone and Morphine Using Ultrafiltration.

AUTHOR: Leow K P; Wright A W E; Cromond T; Smith M T

LOCATION: Brisbane, Australia

SOURCE: Ther.Drug Monit. (15, No. 5, 440-47, 1993) 6 Tab. 23 Ref.

CODEN: TDMODV ISSN: 0163-4356

AVAIL. OF DOC.: Department of Pharmacy, University of Queensland, Brisbane, Queensland 4072, Australia.

LANGUAGE: English

DOCUMENT TYPE: Journal

FIELD AVAIL.: AB; LA; CT; MPC

FILE SEGMENT: Literature

AN 1993-52909 DRUGU P Full-text

AB Serum protein binding of both oxycodone (OX) and morphine (MO) was fairly low and independent of drug concentration in the therapeutic range, but increased with increasing levels of total protein and of purified HSA or human alpha1-acid glycoprotein (AAG, both Sigma-Chemical), in blood samples from healthy subjects. Albumin was the major binding protein for both OX and MO. Bound

ABEX subjects. Albumin was the major binding protein for both OX and MO. Bound

AB respectively. (SAB)

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fractions also increased with a decrease in temperature, and decreased with a small reduction in pH. Serum samples spiked with known OX concentrations showed a gradual decline in serum protein binding with storage time. It is concluded that disease states altering protein concentrations may affect serum protein binding of OX or MO, but that this is unlikely to alter pharmacological effects due to their normally low extent of protein binding.

ABEX Mean (and range) concentrations of total protein, albumin and AAG in blood from healthy volunteers were 74 (62-80), 44 (35-48) and 0.91 (0.55-2.4) g/l, respectively. At physiological pH and temperature, mean serum protein binding (measured by ultrafiltration) was 45.1% for OX and 35.3% for MO. Total and unbound MO and OX were measured by HPLC. A decrease in temperature from 37 to 23 deg. increased serum protein binding by 8-9% for OX and 7-10% for MO. A reduction in pH from 7.75-8.05 to 7.4 reduced serum protein binding by 4-5% for OX and 4-7% for MO. For each pH and temperature variation, serum protein binding for MO was lower than for OX and independent of drug concentration from 5 to 100 ng/ml. Storage of serum samples containing known concentrations of OX at -20 deg resulted in a decline in serum protein binding of OX from about 45% to 39% at 4 wk. Albumin was the major binding protein for both OX and MO, with AAG accounting for only a small proportion of total binding. The bound fraction of OX and MO increased with increasing albumin and AAG concentrations, with higher binding for OX than MO. A reduction in pH to 7.4 and increase in temperature from 23 to 37 deg reduced the binding affinities (Ka) of OX and MO in serum. At each pH and temperature, Ka for MO was lower than that for OX. Binding affinities were higher for AAG than HSA for both OX and MO, and did not change with different protein concentrations. For both OX and MO, Ka was inversely proportional to HSA concentration. The fraction of OX bound to AAG increased with AAG concentration, but the % MO bound was only weakly correlated with AAG concentration. (W103/KP)

L212 ANSWER 8 OF 16 DRUGU COPYRIGHT 2006 THE THOMSON CORP on STN

ACCESSION NUMBER: 1994-06523 DRUGU P Full-text

TITLE: Antinociceptive Potencies of Oxycodone (OC), Noroxycodone (NOC) and Morphine (M) After ICV Administration to Rats.

AUTHOR: Smith M T; Leow K P

CORPORATE SOURCE: Univ.Queensland

LOCATION: Brisbane, Australia

SOURCE: Clin.Expt.Pharmacol.Physiol. (Suppl. 1, 67, 1993) 1 Ref.

CODEN: CEXPB9 ISBN: 0305-1870

AVAIL. OF DOC.: Department of Pharmacy, University of Queensland, Qld 4072, Australia.

LANGUAGE: English

DOCUMENT TYPE: Journal

FIELD AVAIL.: AB; LA; CT

FILE SEGMENT: Literature

AN 1994-06523 DRUGU P Full-text

AB The antinociceptive potencies of oxycodone (OC), its metabolite noroxycodone (NOC) and of morphine (M) were compared after i.c.v. administration in rats, using the tail flick latency test. Analgesic activity was demonstrated for NOC, but it was not as potent as M or OC. I.c.v. naloxone (NAL) blocked the antinociceptive effects of OC and NOC, but failed to totally eliminate the excitatory effects (allodynia, excessive grooming, tremor, Straub tail, myoclonus, etc.) elicited by i.c.v. NOC. It is thus possible that non-opioid mechanisms are involved in the excitatory effects of NOC as has been reported previously for high-dose i.c.v. M. (congress abstract).

ABEX The i.c.v. ED50 values for M, OC, and NOC were 11, 27.5 and 57 ug, respectively. Whilst antinociceptive (more than 50% of maximal possible effect) was noted for all doses of M tested (5 ug or higher), antinociceptive activity was noted only in rats receiving at least 40 ug

AB OC has low affinity for mu-opioid receptor (Ka more than 1 uM), it is postulated that OC's analgesic efficacy may be due to formation of 1 or more active metabolites. The current studies both in cancer patients receiving OC chronically and in healthy volunteers after a single oral dose (10 mg) showed that the mean total urinary recovery of OC, NOC and OM (conjugated and unconjugated) was only 25%. Also, the OM urinary concentration (conjugated and unconjugated) was below the limits of detection (less than 0.5 ug/ml) in all urine samples. After incubation of OC urine with beta-glucuronidase, a new metabolite accounting for at least 50% of the OC dose, appeared in the HPLC chromatogram. In healthy volunteers this new metabolite only appeared in the 2nd 12 hr period after dosing. This putative OC metabolite was difficult to isolate because it is unstable both in unbuffered urine and in HPLC mobile phase. One possible structure for the putative new metabolite of OC which is consistent with the UV spectrum and with its instability in aqueous fluids is a catechol derivative of OM. (ES4/RSV)

L212 ANSWER 9 OF 16 DRUGU COPYRIGHT 2006 THE THOMSON CORP on STN

ACCESSION NUMBER: 1994-06518 DRUGU P Full-text

TITLE: A New Metabolite of Oxycodone in Humans.

AUTHOR: Ross F B; Cromond T; Smith M T

CORPORATE SOURCE: Univ.Queensland

LOCATION: Brisbane, Australia

SOURCE: Clin.Expt.Pharmacol.Physiol. (Suppl. 1, 63, 1993) 1 Ref.

CODEN: CEXPB9 ISBN: 0305-1870

AVAIL. OF DOC.: Department of Pharmacy, University of Queensland, Qld 4072, Australia.

LANGUAGE: English

DOCUMENT TYPE: Journal

FIELD AVAIL.: AB; LA; CT; MPC

FILE SEGMENT: Literature

AN 1994-06518 DRUGU P Full-text

AB The metabolism of oxycodone (OC), a semi-synthetic opioid derivative with a reported efficacy approximately 0.7 that of morphine for the management of cancer pain, was studied in 8 cancer patients receiving OC chronically and in 5 healthy volunteers after a single p.o. dose. Urinary recovery of OC, noroxycodone (NOC) and oxymorphone (OM) was only 25%. However, an unstable metabolite was found, that accounted for at least 50% of the OC dose, and was thought to be a catechol derivative of OM. (congress abstract).

ABEX As OC has low affinity for mu-opioid receptor (Ka more than 1 uM), it is postulated that OC's analgesic efficacy may be due to formation of 1 or more active metabolites. The current studies both in cancer patients receiving OC chronically and in healthy volunteers after a single oral dose (10 mg) showed that the mean total urinary recovery of OC, NOC and OM (conjugated and unconjugated) was only 25%. Also, the OM urinary concentration (conjugated and unconjugated) was below the limits of detection (less than 0.5 ug/ml) in all urine samples. After incubation of OC urine with beta-glucuronidase, a new metabolite accounting for at least 50% of the OC dose, appeared in the HPLC chromatogram. In healthy volunteers this new metabolite only appeared in the 2nd 12 hr period after dosing. This putative OC metabolite was difficult to isolate because it is unstable both in unbuffered urine and in HPLC mobile phase. One possible structure for the putative new metabolite of OC which is consistent with the UV spectrum and with its instability in aqueous fluids is a catechol derivative of OM. (ES4/RSV)

L212 ANSWER 10 OF 16 CAPLUS COPYRIGHT 2006 ACS on STN DUPLICATE 1

ACCESSION NUMBER: 2005-219720 CAPLUS Full-text

DOCUMENT NUMBER: 142-274052

TITLE: Methods and compositions using sub-analgesic doses of a μ opioid agonist and oxycodone for reducing the risk associated with the administration of opioid analgesics in patients with diagnosed or undiagnosed respiratory illness

INVENTOR(S): Pace, Gary W.; Smith, Maree T.

PATENT ASSIGNEE(S): USA

SOURCE: U.S. Pat. Appl. Publ., 11 pp.

DOCUMENT TYPE: Patent

LANGUAGE: English

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
US 2005053659	A1	20050310	US 2003-661458	20030910
WO 2005205621	A1	20050324	WO 2004-US29731	20040910
M: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BW, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, EG, ES, FI, GB, GD, GE, GH, GR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MM, MW, MX, MZ, NA, NI, NO, NZ, OM, PH, PL, PT, RO, RU, SC, SD, SE, SG, SK, SL, SY, TJ, TM, TN, TR, TT, TZ, UA, UG, UZ, VC, VN, YU, ZA, ZM, ZW, RW: BW, GH, KE, LS, MM, MW, NA, SD, SL, SZ, TZ, UG, ZM, ZW, AM, AZ, BY, KQ, KZ, MD, RU, RU, TJ, TM, AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, SI, FI, RO, CY, TR, BG, CZ, EE, HU, PL, SK				
PRIORITY APPLN. INFO.: US 2003-661458 A1 20030910 WO 2004-US29731 W 20040910				

ED Entered STN: 11 Mar 2005

AB The invention discloses methods for reducing the risk associated with the administration of opioid analgesics in patients diagnosed or undiagnosed with respiratory illness by administering an analgesic composition comprising a sub-analgesic dosage of a μ -opioid agonist selected from morphine, fentanyl, sufentanil, alfentanil, oxymorphone and hydromorphone, or a pharmaceutically acceptable salt thereof, and a sub-analgesic dosage of oxycodone, which is a κ -opioid agonist, or a pharmaceutically acceptable salt thereof.

L212 ANSWER 11 OF 16 CAPLUS COPYRIGHT 2006 ACS on STN DUPLICATE 3

ACCESSION NUMBER: 2003:757712 CAPLUS Full-text

DOCUMENT NUMBER: 139:271069

TITLE: Methods and compositions including nitric oxide donors and opioid analgesics for pain relief

INVENTOR(S): Smith, Maree Therese; Brown, Lindsay; Harvey, Mark Bradford Pullar; Williams, Craig Mckenzie

PATENT ASSIGNEE(S): The University of Queensland, Australia

SOURCE: PCT Int. Appl., 69 pp.

CODEN: PIXXD2

DOCUMENT TYPE: Patent

LANGUAGE: English

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 2003078437	A1	20030925	WO 2003-AU335	20030320
M: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, EG, ES, FI, GB, GD, GE, GH, GR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MM, MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE, SI, SK, TJ, TM, TR, TT, UA, UG, UZ, VC, VN, YU, ZA, ZM, ZW, RW: BE, BG, CH, DE, DK, ES, FI, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, SI, FI, RO, CY, TR, BG, CZ, EE, HU, PL, SK				
PRIORITY APPLN. INFO.: WO 2003-AU335 19961021				

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TZ, UA, UG, US, UZ, VC, VN, YU, ZA, ZM, ZW, TZ, UG, ZM, ZW, AM, AZ, BY, RU: GH, GM, KR, LS, MM, MZ, SD, SL, SZ, TZ, UG, ZM, ZW, AM, AZ, BY, KO, KZ, MD, GR, HU, IE, IT, LU, MC, NL, PT, RO, SE, SI, SK, TR, FI, FR, GB, GR, HU, IE, IT, LU, MC, NL, PT, RO, SE, SI, SK, TR, BP, BJ, CF, CO, CI, CM, GA, GN, GO, GW, ML, MR, NE, SN, TD, TG	
CA 2479094 A1 20030935 CA 2003-2479094 20030320	
AU 2003209850 A1 20030929 AU 2003-209850 20030320	
US 2003211944 A1 20031127 US 2003-393050 20030320	
EP 1495026 A1 20050112 EP 2003-744274 20030320	
R: AT, BE, CH, DE, DK, ES, FR, GR, IT, LI, LU, NL, SE, MC, PT, IE, SI, FI, RO, CY, TR, BG, CZ, EE, HU, PL, SK	
JP 2005524676 T2 20050518 JP 2003-576442 20030320	
CN 1703416 A 20051130 CN 2003-609229 20030320	
PRIORITY APPLN. INFO.: US 2002-366594P P 20020320	
OTHER SOURCE(S): MARPAT 139:271069	
ED Entered STN: 26 Sep 2003	
GI	

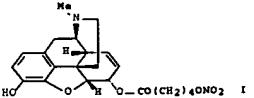
NO 2003078437 A1 20030925 WO 2003-AU335 W 20030320

PRIORITY APPLN. INFO.: US 2003-661458 A1 20030910 WO 2004-US29731 W 20040910

ED Entered STN: 26 Sep 2003

GI

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AB Comps. and methods that induce, promote or otherwise facilitate pain relief are disclosed. These comps. and methods comprise a nitric oxide donor which either directly or indirectly prevents, attenuates or reverses the development of reduced opioid sensitivity, together with a compound which activates the opioid receptor that is the subject of the reduced opioid sensitivity. The comps. and methods prevent or alleviate pain, especially in neuropathic conditions and even more especially in peripheral neuropathic conditions such as painful diabetic neuropathy. The preferred nitric oxide donor is L-arginine, while the preferred comps. which activate the opioid receptor are morphine and oxycodone. Conjugate comps. comprising the nitric oxide donor and an opioid analgesic are also disclosed. Preparation of morphine-NO donor conjugates, e.g. I, is also described.

REFERENCE COUNT: 9 THERE ARE 9 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L212 ANSWER 12 OF 16 CAPLUS COPYRIGHT 2006 ACS on STN DUPLICATE 5

ACCESSION NUMBER: 139:261742 CAPLUS Full-text

DOCUMENT NUMBER: 126:325531

TITLE: Production of analgesic synergy by co-administration of sub-analgesic doses of a μ -opioid agonist and a κ -opioid agonist

INVENTOR(S): Smith, Maree; Ross, Fraser

PATENT ASSIGNEE(S): University of Queensland, Australia; Lynx Project

SOURCE: Limited; Smith, Maree; Ross, Fraser

PCT Int. Appl., 69 pp.

CODEN: PIXXD2

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
ZA 9608808 A 19970527 ZA 1996-8808	19961019			
CA 2235375 AA 19970424 CA 1996-2235375	19961021			
AU 9672076 A1 19970507 AU 1996-72076	19961021			
AU 706691 B2 19990624	19990624			
EP 871488 A1 19981021	19981021			
EP 871488 B1 20050413	20050413			
R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, FI, PH, PL, PT, RO, RU, SC, SD, SE, SG, SK, SL, TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VC, VN, YU, ZA, ZM, ZW, RU: GH, GM, KR, LS, MM, MZ, SD, SL, SZ, TZ, UG, ZM, ZW, AM, AZ, BY, KG, KZ, MD, GR, HU, IE, IT, LU, MC, NL, PT, RO, SE, SI, SK, TR, FI, FR, GB, GR, HU, IE, IT, LU, MC, NL, PT, RO, SE, SI, SK, TR, BP, BJ, CF, CG, CI, CM, GA, GN, GO, GW, ML, MR, NE, SN, TD, TG				
AU 2003209851 A1 20030929 AU 2003-209851 20030320				
US 2003199424 A1 20031023 US 2003-393056 20030320				
PRIORITY APPLN. INFO.: WO 2003-AU336 W 20030320				

ED Entered STN: 26 Sep 2003

AB The invention is involves the use of angiotensin II receptor 1 (AT1 receptor) antagonists for the treatment, prophylaxis, reversal and/or symptomatic relief of a neuropathic condition, especially a peripheral neuropathic condition such as painful diabetic neuropathy, in vertebrate animals and particularly in human subjects. The invention also discloses the use of AT1 receptor antagonists for preventing, attenuating or reversing the development of reduced opioid sensitivity, and more particularly reduced opioid analgesic sensitivity, in individuals and especially in individuals having, or at risk of developing, a neuropathic condition.

REFERENCE COUNT: 4 THERE ARE 4 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L212 ANSWER 14 OF 16 WPIX COPYRIGHT 2006 THE THOMSON CORP on STN

ACCESSION NUMBER: 2006-464147 [47] WPIX

DOC. NO. CPI: C2006-145568 [47]

TITLE: Method of producing analgesia, useful to relieve pain e.g. moderate to severe cancer pain and post surgical pain, comprises administering a nitric oxide donor and an opioid analgesic

DERMINT CLASS: B02

INVENTOR: SMITH M T

PATENT ASSIGNEE: (UYQU-C) UNIV QUEENSLAND

COUNTRY COUNT: 111

PATENT INFO ABBR.:

PATENT NO.	KIND	DATE	WEEK	LA	PG	MAIN IPC
WO 2006066362	A1	20060629	(200647)	87	[1]	

APPLICATION DETAILS:

PATENT NO.	KIND	APPLICATION	DATE
WO 2006066362	A1	WO 2005-AU1976	20051223

PRIORITY APPLN. INFO: AU 2004-907352 20041224

ED 20060724

AN 2006-464147 [47] WPIX

AB WO 2006066362 A1 UPAB: 20060724

NOVELTY - Method of producing analgesia (A) in a subject comprises administering a nitric oxide donor (II) and an opioid analgesic, where (II) delivers nitric oxide (III) at a rate of 0.0002-2 nmol/kg/hour.

DETAILED DESCRIPTION - An INDEPENDENT CLAIM is also included for new compounds of formula (I).

ACTIVITY - Analgesic.

MECHANISM OF ACTION - None given.

USE - (A) is useful to relieve pain (moderate to severe cancer pain, moderate to severe post surgical pain, pain following physical trauma, pain associated

DOCUMENT TYPE: Patent

LANGUAGE: English

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
NO 9714438 A1 19970424	WO 1996-AU656	19961021		
M: AE, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, EG, ES, FI, GB, GD, GE, GH, GM, HR, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MM, MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE, SI, SK, TJ, TM, TR, TT, UA, UG, UZ, VC, VN, YU, ZA, ZM, ZW, RW: BE, BG, CH, DE, DK, ES, FI, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, SI, FI, RO, CY, TR, BG, CZ, EE, HU, PL, SK				
PRIORITY APPLN. INFO.: WO 1996-AU656 19961021				

ED Entered STN: 11 Jun 1997

AB An analgesic composition comprises a sub-analgesic dosage of a μ -opioid agonist or analog or derivative or pharmaceutically acceptable salts thereof and a sub-analgesic dosage of a κ -2-opioid agonist or analog or derivative or pharmaceutically acceptable salts thereof. The μ -opioid agonist may be morphine, fentanyl, sufentanil, alfentanil, or hydromorphone; the κ -2-opioid agonist may be oxycodone.

L212 ANSWER 13 OF 16 CAPLUS COPYRIGHT 2006 ACS on STN

ACCESSION NUMBER: 2003:757520 CAPLUS Full-text

DOCUMENT NUMBER: 139:255390

TITLE: Method of treatment and prophylaxis of neuropathic condition

INVENTOR(S): Smith, Maree Therese; Brown, Lindsay

PATENT ASSIGNEE(S): The University of Queensland, Australia

SOURCE: PCT Int. Appl., 87 pp.

DOCUMENT TYPE: Patent

LANGUAGE: English

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 2003077912 A1	20030925	WO 2003-AU336	20030320	
M: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, EG, ES, FI, GB, GD, GE, GH, GR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MM, MW, MX, MZ, NI, NO, NZ, OM, PH, PL, PT, RO, RU, SC, SD, SE, SG, SK, SL, TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VC, VN, YU, ZA, ZM, ZW, RW: BE, BG, CH, DE, DK, ES, FI, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, SI, FI, RO, CY, TR, BG, CZ, EE, HU, PL, SK				
PRIORITY APPLN. INFO.: WO 2003-AU336 20030320				

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with cardiac infarction and inflammatory pain) (claimed). No biological data given.
ADVANTAGE - (A) enhances the endogenous production of nitrosothiols and reduces the endogenous production of peroxynitrite.

L212 ANSWER 15 OF 16 EMBASE COPYRIGHT (c) 2006 Elsevier B.V. All rights reserved on STN

ACCESSION NUMBER: 2005175148 EMBASE Full-text
TITLE: Co-administration of oxycodone and morphine and analgesic synergy re-examined [1] (multiple letters).
AUTHOR: Smith M.T.; De La Iglesia F.A.; Grath M.; Massalha M.; Pud D.; Adler R.; Eisenberg E.
CORPORATE SOURCE: F.A. De La Iglesia, University of Michigan, Medical School, Ann Arbor, MI, Australia. delegif@umich.edu
SOURCE: British Journal of Clinical Pharmacology, (2005) Vol. 59, NO. 4, pp. 456-488.
ISBN: 0306-5251 CODEN: BCPHEM
COUNTRY: United Kingdom
DOCUMENT TYPE: Journal; Letter
FILE SEGMENT: 008 Neurology and Neurosurgery
037 Drug Literature Index
LANGUAGE: English
ENTRY DATE: Entered STN: 12 May 2005
Last Updated on STN: 12 May 2005
ED Entered STN: 12 May 2005
Last Updated on STN: 12 May 2005
DATA NOT AVAILABLE FOR THIS ACCESSION NUMBER

L212 ANSWER 16 OF 16 EMBASE COPYRIGHT (c) 2006 Elsevier B.V. All rights reserved on STN

ACCESSION NUMBER: 97021905 EMBASE Full-text
DOCUMENT NUMBER: 1597021905
TITLE: HIV-1 protease inhibitors: A review for clinicians.
AUTHOR: Deeks S.G.; Smith M.; Holodniy M.; Kahn J.O.
CORPORATE SOURCE: Dr. J.O. Kahn, University of California, San Francisco General Hospital, 995 Potrero Ave, San Francisco, CA 94110, United States. jkahn@sfraids.ucsf.edu
SOURCE: Journal of the American Medical Association, (1997) Vol. 277, No. 2, pp. 145-153.
Refs: 59
ISSN: 0098-7484 CODEN: JAMAAP
COUNTRY: United States
DOCUMENT TYPE: Journal; General Review
FILE SEGMENT: 006 Internal Medicine
030 Pharmacology
037 Drug Literature Index
038 Adverse Reactions Titles
LANGUAGE: English
SUMMARY LANGUAGE: English
ENTRY DATE: Entered STN: 15 Feb 1997
Last Updated on STN: 15 Feb 1997
ED Entered STN: 15 Feb 1997
Last Updated on STN: 15 Feb 1997

AB Objective.-The clinical care of people infected with human immunodeficiency virus (HIV) has been substantially affected by the introduction of HIV-specific protease inhibitors (PIs). The 4 PIs available are saquinavir mesylate, ritonavir, indinavir sulfate, and nelfinavir mesylate. Comparison studies have not been reported; therefore, an assessment of the available data to aid clinicians and patients in choosing appropriate treatment will be

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presented. Data Sources.-A systematic review of peer-reviewed publications, abstracts from national and international conferences, and product registration information through September 1996. Study Selection and Data Extraction.-Criteria used to select studies include their relevance to PIs, having been published in the English language, and pertinence for clinicians. Data quality and validity included the venue of the publication and relevance to clinical care. Data Synthesis.-Oral administration of ritonavir, indinavir, or nelfinavir generates sustainable drug serum levels to effectively inhibit the protease enzyme; however, saquinavir may not generate sustained levels necessary to inhibit the protease enzyme. Patients treated with ritonavir, indinavir, or nelfinavir experience similar reductions in viral load and increases in CD4+ lymphocytes; smaller effects occur among those treated with saquinavir. Two randomized placebo-controlled studies conducted among patients with severe immune system suppression and substantial sideeffects treatment experience demonstrated reduced HIV disease progression and reduced mortality with PI treatment. Genotypic resistance to PIs occurs; the clinical relevance of resistance is unclear. The costs of these agents including required monitoring impose new and substantial costs. Conclusions.-The PIs have emerged as critical drugs for people with HIV infection. Optimal use involves combination with reverse transcriptase inhibitors. Resistance develops to each agent, and cross-resistance is likely. These agents must be used at full doses with attention to ensuring patient compliance. The expense of these agents may be offset by forestalling disease progression and death and returning people to productive life. Selecting the initial PI must be individualized, and factors to consider include proven activity, possible toxicities, dosing regimens, drug interactions, and costs.

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FILE LAST UPDATED: 13 Dec 2006 (20061213/ED)

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L6 1 SEA FILE=REGISTRY ABB=ON FENTANYL/CN
L7 1 SEA FILE=REGISTRY ABB=ON SUFENTANIL/CN
L8 1 SEA FILE=REGISTRY ABB=ON ALFENTANIL/CN
L9 1 SEA FILE=REGISTRY ABB=ON OXYMORPHONE/CN
L10 1 SEA FILE=REGISTRY ABB=ON HYDROMORPHONE/CN
L11 1 SEA FILE=REGISTRY ABB=ON OXYCODEONE/CN
L12 31087 SEA FILE=CAPLUS ABB=ON (L5 OR L6 OR L7 OR L8 OR L9 OR L10)
L13 1073 SEA FILE=CAPLUS ABB=ON L11
L14 12914 SEA FILE=CAPLUS ABB=ON OPIOIDS/CT
L15 1209 SEA FILE=CAPLUS ABB=ON L14(L)KAPPA/OBI
L16 1944 SEA FILE=CAPLUS ABB=ON L14(L)MU/OBI
L17 56591 SEA FILE=CAPLUS ABB=ON AGONISTS/OBI
L18 368 SEA FILE=CAPLUS ABB=ON L15(L)L17
L19 454 SEA FILE=CAPLUS ABB=ON L16(L)L17
L20 19117 SEA FILE=CAPLUS ABB=ON RESPIRATORY TRACT/OBI
L21 76 SEA FILE=CAPLUS ABB=ON L20(L)CARCINOMA/OBI
L22 25232 SEA FILE=CAPLUS ABB=ON ASTHMA/OBI
L23 424 SEA FILE=CAPLUS ABB=ON BRONCHIECTASIS/OBI OR BRONCHITIS/OBI (L)DI
L24 28786 SEA FILE=CAPLUS ABB=ON LATATION/OBI OR KARTAGENERI/OBI
L25 4089 SEA FILE=CAPLUS ABB=ON TUBERCULOSIS/OBI
L26 120 SEA FILE=CAPLUS ABB=ON RESPIRATORY SYSTEM, NEOPLASM/CT
L27 35560 SEA FILE=CAPLUS ABB=ON LUNG, NEOPLASM/CT
L28 4982 SEA FILE=CAPLUS ABB=ON CHRONIC OBSTRUCTIVE PULMONARY/OBI OR
COPD/OBI
L29 7726 SEA FILE=CAPLUS ABB=ON BRONCHOPNEUMONIA/OBI OR PNEUMONIA/OBI
L30 136 SEA FILE=CAPLUS ABB=ON LARYNgeITIS/OBI
L31 1101 SEA FILE=CAPLUS ABB=ON SINUSITIS/OBI
L32 2601 SEA FILE=CAPLUS ABB=ON EMPHYSEMA/OBI
L33 6378 SEA FILE=CAPLUS ABB=ON FIBROSING/OBI(L)ALVEOLITIS/OBI OR
(PULMONARY/OBI OR LUNG/OBI OR RESPIRATORY/OBI) (L) (FIBROSIS/OBI
OR SARCOIDOSIS/OBI)
L34 6 SEA FILE=CAPLUS ABB=ON SLEEP DISORDERS/CT(L)RESPIRATORY/OBI
L35 943 SEA FILE=CAPLUS ABB=ON SLEEP/OBI(L)APNEA/OBI
L36 1691 SEA FILE=CAPLUS ABB=ON SARCOIDOSIS/CT
L37 39125 SEA FILE=CAPLUS ABB=ON DRUG INTERACTIONS+OLD,NT/CT
L38 4450 SEA FILE=CAPLUS ABB=ON DRUG DELIVERY SYSTEMS+OLD/CT(L)COMB?/OB

(PULMONARY/OBI OR LUNG/OBI OR RESPIRATORY/OBI) (L) (FIBROSIS/OBI
OR SARCOIDOSIS/OBI)
L39 16989 SEA FILE=CAPLUS ABB=ON COMBINATION CHEMOTHERAPY/CT
L40 5480 SEA FILE=CAPLUS ABB=ON COMB?/OBI(L)PHARMAC?/OBI
L41 6 SEA FILE=CAPLUS ABB=ON (L12 OR L19) AND (L13 OR L18) AND (L21
OR L22 OR L23 OR L24 OR L25 OR L26 OR L27 OR L28 OR L29 OR L30
OR L31 OR L32 OR L33 OR L34 OR L35 OR L36) AND (L37 OR L38 OR
L39 OR L40)
L5 1 SEA FILE=REGISTRY ABB=ON MORPHINE/CN
L6 1 SEA FILE=REGISTRY ABB=ON FENTANYL/CN
L7 1 SEA FILE=REGISTRY ABB=ON SUFENTANIL/CN
L8 1 SEA FILE=REGISTRY ABB=ON ALFENTANIL/CN
L9 1 SEA FILE=REGISTRY ABB=ON OXYMORPHONE/CN
L10 1 SEA FILE=REGISTRY ABB=ON HYDROMORPHONE/CN
L11 1 SEA FILE=REGISTRY ABB=ON OXYCODEONE/CN
L12 31087 SEA FILE=CAPLUS ABB=ON (L5 OR L6 OR L7 OR L8 OR L9 OR L10)
L13 1073 SEA FILE=CAPLUS ABB=ON L11
L14 12914 SEA FILE=CAPLUS ABB=ON OPIOIDS/CT
L15 1209 SEA FILE=CAPLUS ABB=ON L14(L)KAPPA/OBI
L16 1944 SEA FILE=CAPLUS ABB=ON L14(L)MU/OBI
L17 56591 SEA FILE=CAPLUS ABB=ON AGONISTS/OBI
L18 368 SEA FILE=CAPLUS ABB=ON L15(L)L17
L19 454 SEA FILE=CAPLUS ABB=ON L16(L)L17
L20 19117 SEA FILE=CAPLUS ABB=ON RESPIRATORY TRACT/OBI
L21 76 SEA FILE=CAPLUS ABB=ON L20(L)CARCINOMA/OBI
L22 25232 SEA FILE=CAPLUS ABB=ON ASTHMA/OBI
L23 424 SEA FILE=CAPLUS ABB=ON BRONCHIECTASIS/OBI OR BRONCHITIS/OBI (L)DI
L24 28786 SEA FILE=CAPLUS ABB=ON LATATION/OBI OR KARTAGENERI/OBI
L25 4089 SEA FILE=CAPLUS ABB=ON TUBERCULOSIS/OBI
L26 120 SEA FILE=CAPLUS ABB=ON RESPIRATORY SYSTEM, NEOPLASM/CT
L27 35560 SEA FILE=CAPLUS ABB=ON LUNG, NEOPLASM/CT
L28 4982 SEA FILE=CAPLUS ABB=ON CHRONIC OBSTRUCTIVE PULMONARY/OBI OR
COPD/OBI
L29 7726 SEA FILE=CAPLUS ABB=ON BRONCHOPNEUMONIA/OBI OR PNEUMONIA/OBI
L30 136 SEA FILE=CAPLUS ABB=ON LARYNgeITIS/OBI
L31 1101 SEA FILE=CAPLUS ABB=ON SINUSITIS/OBI
L32 2601 SEA FILE=CAPLUS ABB=ON EMPHYSEMA/OBI
L33 6378 SEA FILE=CAPLUS ABB=ON FIBROSING/OBI(L)ALVEOLITIS/OBI OR
(PULMONARY/OBI OR LUNG/OBI OR RESPIRATORY/OBI) (L) (FIBROSIS/OBI
OR SARCOIDOSIS/OBI)
L34 6 SEA FILE=CAPLUS ABB=ON SLEEP DISORDERS/CT(L)RESPIRATORY/OBI
L35 943 SEA FILE=CAPLUS ABB=ON SLEEP/OBI(L)APNEA/OBI
L36 1691 SEA FILE=CAPLUS ABB=ON SARCOIDOSIS/CT
L37 552 SEA FILE=CAPLUS ABB=ON (L12 OR L19) (L) (COMB?/OBI OR COADMIN?/OBI
OR CODRUG?/OBI OR CONCOMITANT?/OBI OR CONCURRENT?/OBI OR
BLEND?/OBI OR MIXTURE?/OBI)
L38 82 SEA FILE=CAPLUS ABB=ON (L13 OR L18) (L) (COMB?/OBI OR COADMIN?/OBI
OR CODRUG?/OBI OR CONCOMITANT?/OBI OR CONCURRENT?/OBI OR
BLEND?/OBI OR MIXTURE?/OBI)
L39 3 SEA FILE=CAPLUS ABB=ON L42 AND L43 AND (L21 OR L22 OR L23 OR

L24 OR L25 OR L26 OR L27 OR L28 OR L29 OR L30 OR L31 OR L32 OR
L33 OR L34 OR L35 OR L36)

>> = 141,144 not 1210

L213 5 (L41 OR L44) NOT L210

>> fil embase; d que 182; d que 184

FILE 'EMBASE' ENTERED AT 11:12:33 ON 14 DEC 2006

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FILE COVERS 1974 TO 13 Dec 2006 (20061213/ED)

EMBASE is now updated daily. SDI frequency remains weekly (default)
and biweekly.

This file contains CAS Registry Numbers for easy and accurate
substance identification.

L48 53452 SEA FILE=EMBASE ABB=ON MORPHINE/CT
L49 26736 SEA FILE=EMBASE ABB=ON FENTANYL/CT OR FENTANYL CITRATE/CT
L50 4395 SEA FILE=EMBASE ABB=ON SUFENTANIL/CT OR SUFENTANIL CITRATE/CT

L51 4482 SEA FILE=EMBASE ABB=ON ALFENTANIL/CT

L52 805 SEA FILE=EMBASE ABB=ON OXYMORPHONE/CT

L53 2957 SEA FILE=EMBASE ABB=ON HYDROMORPHONE/CT

L54 3754 SEA FILE=EMBASE ABB=ON OXYCODEONE/CT

L55 84233 SEA FILE=EMBASE ABB=ON ASTHMA-/NT/CT

L56 4535 SEA FILE=EMBASE ABB=ON BRONCHIECTASIS-/NT/CT

L57 15140 SEA FILE=EMBASE ABB=ON LUNG TUBERCULOSIS/CT

L58 26377 SEA FILE=EMBASE ABB=ON CHRONIC OBSTRUCTIVE LUNG DISEASE/CT

L59 22047 SEA FILE=EMBASE ABB=ON BRONCHITIS-/NT/CT

L60 2275 SEA FILE=EMBASE ABB=ON LUNG TUBERCULOSIS/CT

L61 2500 SEA FILE=EMBASE ABB=ON LARYNGITIS-/NT/CT

L62 12991 SEA FILE=EMBASE ABB=ON SINUSITIS-/NT/CT

L63 13249 SEA FILE=EMBASE ABB=ON EMPHYSEMA-/NT/CT

L64 2738 SEA FILE=EMBASE ABB=ON FIBROSING ALVEOLITIS/CT

L65 19527 SEA FILE=EMBASE ABB=ON LUNG FIBROSIS-/NT/CT

L66 11397 SEA FILE=EMBASE ABB=ON SARCOIDOSIS/CT

L67 91685 SEA FILE=EMBASE ABB=ON LUNG CANCER-/NT/CT

L68 11977 SEA FILE=EMBASE ABB=ON SLEEP APNEA SYNDROME/CT

L69 38068 SEA FILE=EMBASE ABB=ON DRUG POTENTIATION/CT

L70 1228 SEA FILE=EMBASE ABB=ON MU OPIATE RECEPTOR AGONIST/CT

L71 949 SEA FILE=EMBASE ABB=ON KAPPA OPIATE RECEPTOR AGONIST/CT

L72 0 SEA FILE=EMBASE ABB=ON (L48 OR L49 OR L50 OR L51 OR L52 OR

L53 OR L76) AND (L77 OR L54) AND L75 AND (L55 OR L56 OR L57 OR

L58 OR L59 OR L60 OR L61 OR L62 OR L63 OR L64 OR L65 OR L66 OR

L67 OR L68)

L73 53452 SEA FILE=EMBASE ABB=ON MORPHINE/CT

L74 26736 SEA FILE=EMBASE ABB=ON FENTANYL/CT OR FENTANYL CITRATE/CT

L75 4395 SEA FILE=EMBASE ABB=ON SUFENTANIL/CT OR SUFENTANIL CITRATE/CT

L76 4482 SEA FILE=EMBASE ABB=ON ALFENTANIL/CT

L77 805 SEA FILE=EMBASE ABB=ON OXYMORPHONE/CT

L78 11977 SEA FILE=EMBASE ABB=ON MORPHINS/CN

L79 1 SEA FILE=REGISTRY ABB=ON FENTANYL/CN

L80 1 SEA FILE=REGISTRY ABB=ON SUFENTANIL/CN

L81 1 SEA FILE=REGISTRY ABB=ON ALFENTANIL/CN

L82 1 SEA FILE=REGISTRY ABB=ON OXYMORPHONE/CN

L83 1 SEA FILE=REGISTRY ABB=ON HYDROMORPHONE/CN

L84 1 SEA FILE=REGISTRY ABB=ON OXYCODEONE/CN

L85 26377 SEA FILE=EMBASE ABB=ON (L72 OR L79) AND (L80 OR L78) AND (L55

OR L56 OR L57 OR L58 OR L59 OR L60 OR L61 OR L62 OR L63 OR L64

OR L65 OR L66 OR L67 OR L68)

L53 2957 SEA FILE=EMBASE ABB=ON HYDROMORPHONE/CT

L54 3754 SEA FILE=EMBASE ABB=ON OXYCODEONE/CT

L55 84233 SEA FILE=EMBASE ABB=ON ASTHMA-/NT/CT

L56 4535 SEA FILE=EMBASE ABB=ON BRONCHIECTASIS-/NT/CT

L57 15140 SEA FILE=EMBASE ABB=ON LUNG TUBERCULOSIS/CT

L58 26377 SEA FILE=EMBASE ABB=ON CHRONIC OBSTRUCTIVE LUNG DISEASE/CT

L59 22047 SEA FILE=EMBASE ABB=ON BRONCHITIS-/NT/CT

L60 2275 SEA FILE=EMBASE ABB=ON LUNG TUBERCULOSIS/CT

L61 2500 SEA FILE=EMBASE ABB=ON LARYNGITIS-/NT/CT

L62 12991 SEA FILE=EMBASE ABB=ON SINUSITIS-/NT/CT

L63 13249 SEA FILE=EMBASE ABB=ON EMPHYSEMA-/NT/CT

L64 2738 SEA FILE=EMBASE ABB=ON FIBROSING ALVEOLITIS/CT

L65 19527 SEA FILE=EMBASE ABB=ON LUNG FIBROSIS-/NT/CT

L66 11397 SEA FILE=EMBASE ABB=ON SARCOIDOSIS/CT

L67 91685 SEA FILE=EMBASE ABB=ON LUNG CANCER-/NT/CT

L68 11977 SEA FILE=EMBASE ABB=ON SLEEP APNEA SYNDROME/CT

L69 38068 SEA FILE=EMBASE ABB=ON DRUG POTENTIATION/CT

L70 1228 SEA FILE=EMBASE ABB=ON MU OPIATE RECEPTOR AGONIST/CT

L71 949 SEA FILE=EMBASE ABB=ON KAPPA OPIATE RECEPTOR AGONIST/CT

L72 0 SEA FILE=EMBASE ABB=ON (L48 OR L49 OR L50 OR L51 OR L52 OR

L53 OR L76) AND (L77 OR L54) AND L75 AND (L55 OR L56 OR L57 OR

L58 OR L59 OR L60 OR L61 OR L62 OR L63 OR L64 OR L65 OR L66 OR

L67 OR L68)

L73 53452 SEA FILE=EMBASE ABB=ON MORPHINE/CT

L74 26736 SEA FILE=EMBASE ABB=ON FENTANYL/CT OR FENTANYL CITRATE/CT

L75 4395 SEA FILE=EMBASE ABB=ON SUFENTANIL/CT OR SUFENTANIL CITRATE/CT

L76 4482 SEA FILE=EMBASE ABB=ON ALFENTANIL/CT

L77 805 SEA FILE=EMBASE ABB=ON OXYMORPHONE/CT

L78 11977 SEA FILE=EMBASE ABB=ON SLEEP APNEA SYNDROME/CT

L79 38068 SEA FILE=EMBASE ABB=ON DRUG POTENTIATION/CT

L80 1228 SEA FILE=EMBASE ABB=ON MU OPIATE RECEPTOR AGONIST/CT

L81 949 SEA FILE=EMBASE ABB=ON KAPPA OPIATE RECEPTOR AGONIST/CT

L82 0 SEA FILE=EMBASE ABB=ON (L48 OR L49 OR L50 OR L51 OR L52 OR

L53 OR L76) AND (L77 OR L54) AND L75 AND (L55 OR L56 OR L57 OR

L58 OR L59 OR L60 OR L61 OR L62 OR L63 OR L64 OR L65 OR L66 OR

L67 OR L68)

L83 53452 SEA FILE=EMBASE ABB=ON MORPHINE/CT

L84 26736 SEA FILE=EMBASE ABB=ON FENTANYL/CT OR FENTANYL CITRATE/CT

L85 4395 SEA FILE=EMBASE ABB=ON SUFENTANIL/CT OR SUFENTANIL CITRATE/CT

L86 4482 SEA FILE=EMBASE ABB=ON ALFENTANIL/CT

L87 805 SEA FILE=EMBASE ABB=ON OXYMORPHONE/CT

L88 11977 SEA FILE=EMBASE ABB=ON SLEEP APNEA SYNDROME/CT

L89 38068 SEA FILE=EMBASE ABB=ON DRUG POTENTIATION/CT

L90 1228 SEA FILE=EMBASE ABB=ON MU OPIATE RECEPTOR AGONIST/CT

L91 949 SEA FILE=EMBASE ABB=ON KAPPA OPIATE RECEPTOR AGONIST/CT

L92 2660 SEA FILE=EMBASE ABB=ON ALFENTANIL/CT

L93 11360 SEA FILE=EMBASE ABB=ON OXYMORPHONE/CT

L94 8806 SEA FILE=EMBASE ABB=ON SLEEP APNEA SYNDROME/CT

L95 3147 SEA FILE=EMBASE ABB=ON SLEEP APNEA SYNDROME/CT

L96 11397 SEA FILE=EMBASE ABB=ON SLEEP APNEA SYNDROME/CT

L97 13249 SEA FILE=EMBASE ABB=ON SLEEP APNEA SYNDROME/CT

L98 13249 SEA FILE=EMBASE ABB=ON SLEEP APNEA SYNDROME/CT

L99 13249 SEA FILE=EMBASE ABB=ON SLEEP APNEA SYNDROME/CT

L100 13249 SEA FILE=EMBASE ABB=ON SLEEP APNEA SYNDROME/CT

L101 13249 SEA FILE=EMBASE ABB=ON SLEEP APNEA SYNDROME/CT

L102 13249 SEA FILE=EMBASE ABB=ON SLEEP APNEA SYNDROME/CT

L103 13249 SEA FILE=EMBASE ABB=ON SLEEP APNEA SYNDROME/CT

L104 13249 SEA FILE=EMBASE ABB=ON SLEEP APNEA SYNDROME/CT

L105 13249 SEA FILE=EMBASE ABB=ON SLEEP APNEA SYNDROME/CT

L106 13249 SEA FILE=EMBASE ABB=ON SLEEP APNEA SYNDROME/CT

L107 13249 SEA FILE=EMBASE ABB=ON SLEEP APNEA SYNDROME/CT

L108 13249 SEA FILE=EMBASE ABB=ON SLEEP APNEA SYNDROME/CT

L109 13249 SEA FILE=EMBASE ABB=ON SLEEP APNEA SYNDROME/CT

L110 13249 SEA FILE=EMBASE ABB=ON SLEEP APNEA SYNDROME/CT

L111 13249 SEA FILE=EMBASE ABB=ON SLEEP APNEA SYNDROME/CT

L112 13249 SEA FILE=EMBASE ABB=ON SLEEP APNEA SYNDROME/CT

L113 13249 SEA FILE=EMBASE ABB=ON SLEEP APNEA SYNDROME/CT

L114 13249 SEA FILE=EMBASE ABB=ON SLEEP APNEA SYNDROME/CT

L115 13249 SEA FILE=EMBASE ABB=ON SLEEP APNEA SYNDROME/CT

L116 13249 SEA FILE=EMBASE ABB=ON SLEEP APNEA SYNDROME/CT

L117 13249 SEA FILE=EMBASE ABB=ON SLEEP APNEA SYNDROME/CT

L118 13249 SEA FILE=EMBASE ABB=ON SLEEP APNEA SYNDROME/CT

L119 13249 SEA FILE=EMBASE ABB=ON SLEEP APNEA SYNDROME/CT

L120 13249 SEA FILE=EMBASE ABB=ON SLEEP APNEA SYNDROME/CT

L121 13249 SEA FILE=EMBASE ABB=ON SLEEP APNEA SYNDROME/CT

L122 13249 SEA FILE=EMBASE ABB=ON SLEEP APNEA SYNDROME/CT

L123 13249 SEA FILE=EMBASE ABB=ON SLEEP APNEA SYNDROME/CT

L124 13249 SEA FILE=EMBASE ABB=ON SLEEP APNEA SYNDROME/CT

L125 13249 SEA FILE=EMBASE ABB=ON SLEEP APNEA SYNDROME/CT

L126 13249 SEA FILE=EMBASE ABB=ON SLEEP APNEA SYNDROME/CT

L127 13249 SEA FILE=EMBASE ABB=ON SLEEP APNEA SYNDROME/CT

L128 13249 SEA FILE=EMBASE ABB=ON SLEEP APNEA SYNDROME/CT

L129 13249 SEA FILE=EMBASE ABB=ON SLEEP APNEA SYNDROME/CT

L130 13249 SEA FILE=EMBASE ABB=ON SLEEP APNEA SYNDROME/CT

L131 13249 SEA FILE=EMBASE ABB=ON SLEEP APNEA SYNDROME/CT

L132 13249 SEA FILE=EMBASE ABB=ON SLEEP APNEA SYNDROME/CT

L133 13249 SEA FILE=EMBASE ABB=ON SLEEP APNEA SYNDROME/CT

L134 13249 SEA FILE=EMBASE ABB=ON SLEEP APNEA SYNDROME/CT

L135 13249 SEA FILE=EMBASE ABB=ON SLEEP APNEA SYNDROME/CT

L136 13249 SEA FILE=EMBASE ABB=ON SLEEP APNEA SYNDROME/CT

L137 13249 SEA FILE=EMBASE ABB=ON SLEEP APNEA SYNDROME/CT

L138 13249 SEA FILE=EMBASE ABB=ON SLEEP APNEA SYNDROME/CT

L139 13249 SEA FILE=EMBASE ABB=ON SLEEP APNEA SYNDROME/CT

L140 13249 SEA FILE=EMBASE ABB=ON SLEEP APNEA SYNDROME/CT

L141 13249 SEA FILE=EMBASE ABB=ON SLEEP APNEA SYNDROME/CT

OR COADMIN?/BI, ABBX OR CONCOMITANT?/BI, ABBX OR CONCURRENT?/BI, ABBX OR BLEND?/BI, ABBX OR MIX?/BI, ABBX)
L142 2 SEA FILE+WPIX ABB-ON L141 AND (L124 OR L125 OR L126 OR L127
OR L128 OR L129)

=> # L134, L142 not L211
L216 2 (L134 OR L142) NOT L211

=> fil medl; d que 1189; d que 1197; d que 1207; d que 1179

FILE 'MEDLINE' ENTERED AT 11:13:43 ON 14 DEC 2006

FILE LAST UPDATED: 13 Dec 2006 (20061213/UP). FILE COVERS 1950 TO DATE.

In preparation for the annual MEDLINE reload, the National Library of Medicine (NLM) has suspended delivery of regular updates as of November 15, 2006. In-process and in-data-review records will resume delivery on November 21, 2006, and will continue to be added to MEDLINE until December 17, 2006.

On December 17, 2006, all regular MEDLINE updates from November 15 to December 16 will be added to MEDLINE, along with 2007 Medical Subject Headings (MeSH(R)) and 2007 tree numbers.

The annual reload will be available in early 2007.

This file contains CAS Registry Numbers for easy and accurate substance identification.

L180(28104)SEA FILE+MEDLINE ABB-ON MORPHINE/CT
L181(10382)SEA FILE+MEDLINE ABB-ON FENTANYL-NT/CT
L182(294)SEA FILE+MEDLINE ABB-ON OXYMORPHONE/CT
L183(704)SEA FILE+MEDLINE ABB-ON HYDROMORPHONE/CT
L184(540)SEA FILE+MEDLINE ABB-ON OXYCODONE/CT
L185(108974)SEA FILE+MEDLINE ABB-ON DRUG INTERACTIONS-NT/CT
L186(42787)SEA FILE+MEDLINE ABB-ON DRUG COMBINATIONS/CT
L187(97253)SEA FILE+MEDLINE ABB-ON DRUG THERAPY, COMBINATION/CT
L188(15)SEA FILE+MEDLINE ABB-ON (L180 OR L181 OR L182 OR L183) AND
L184 AND (L185 OR L186 OR L187)
L189 3 SEA FILE+MEDLINE ABB-ON L188 AND SYNERG?

L190(108974)SEA FILE+MEDLINE ABB-ON DRUG INTERACTIONS-NT/CT
L191(42787)SEA FILE+MEDLINE ABB-ON DRUG COMBINATIONS/CT
L192(97253)SEA FILE+MEDLINE ABB-ON DRUG THERAPY, COMBINATION/CT
L193(1136)SEA FILE+MEDLINE ABB-ON RECEPTORS, OPIOID, MU/CT(L)AG/CT
L194(881)SEA FILE+MEDLINE ABB-ON RECEPTORS, OPIOID, KAPPA/CT(L)AG/CT
L195(23)SEA FILE+MEDLINE ABB-ON L193 AND L194 AND (L190 OR L191 OR
L192)
L196(240557)SEA FILE+MEDLINE ABB-ON DOSE-RESPONSE RELATIONSHIP, DRUG/CT
L197 1 SEA FILE+MEDLINE ABB-ON L195 AND L196 AND CONDITIONING.
OPERANT/CT

L198(28104)SEA FILE+MEDLINE ABB-ON MORPHINE/CT
L199(10382)SEA FILE+MEDLINE ABB-ON FENTANYL-NT/CT
L200(294)SEA FILE+MEDLINE ABB-ON OXYMORPHONE/CT
L201(704)SEA FILE+MEDLINE ABB-ON HYDROMORPHONE/CT
L202(540)SEA FILE+MEDLINE ABB-ON OXYCODONE/CT
L203(1136)SEA FILE+MEDLINE ABB-ON RECEPTORS, OPIOID, MU/CT(L)AG/CT
L204(881)SEA FILE+MEDLINE ABB-ON RECEPTORS, OPIOID, KAPPA/CT(L)AG/CT
L205(488)SEA FILE+MEDLINE ABB-ON (L198 OR L199 OR L200 OR L201 OR
L203) AND (L202 OR L204)
L206(8267)SEA FILE+MEDLINE ABB-ON COUGH/CT
L207 1 SEA FILE+MEDLINE ABB-ON L205 AND L206

L164(28104)SEA FILE+MEDLINE ABB-ON MORPHINE/CT
L165(10382)SEA FILE+MEDLINE ABB-ON FENTANYL-NT/CT
L166(294)SEA FILE+MEDLINE ABB-ON OXYMORPHONE/CT
L167(704)SEA FILE+MEDLINE ABB-ON HYDROMORPHONE/CT
L168(540)SEA FILE+MEDLINE ABB-ON OXYCODONE/CT
L169(124991)SEA FILE+MEDLINE ABB-ON LUNG DISEASES, OBSTRUCTIVE-NT/CT
L170(5936)SEA FILE+MEDLINE ABB-ON BRONCHIECTASIS-NT/CT
L171(57086)SEA FILE+MEDLINE ABB-ON TUBERCULOSIS, PULMONARY-NT/CT
L172(1460)SEA FILE+MEDLINE ABB-ON BRONCHOPNEUMONIA/CT
L173(3610)SEA FILE+MEDLINE ABB-ON LARYNGITIS-NT/CT
L174(11628)SEA FILE+MEDLINE ABB-ON SINUSITIS-NT/CT
L175(13172)SEA FILE+MEDLINE ABB-ON PULMONARY FIBROSIS/CT
L176(1561)SEA FILE+MEDLINE ABB-ON SARCOIDOSIS, PULMONARY/CT
L177(112814)SEA FILE+MEDLINE ABB-ON LUNG NEOPLASMS-NT/CT
L178(12706)SEA FILE+MEDLINE ABB-ON SLEEP APNEA SYNDROMES-NT/CT
L179 1 SEA FILE+MEDLINE ABB-ON (L164 OR L165 OR L166 OR L167) AND
L168 AND (L169 OR L171 OR L172 OR L173 OR L174 OR L175
OR L176 OR L177 OR L178)

=> # L1189, L1197, L1207, L1179

L217 6 (L1189 OR L1197 OR L1207 OR L1179)

=> => dup rem L1217, L1215, L1213, L1216, L1214
FILE 'MEDLINE' ENTERED AT 11:13:15 ON 14 DEC 2006

FILE 'DRUGU' ENTERED AT 11:13:15 ON 14 DEC 2006
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PROCESSING COMPLETED FOR L217
PROCESSING COMPLETED FOR L215
PROCESSING COMPLETED FOR L213
PROCESSING COMPLETED FOR L216
PROCESSING COMPLETED FOR L214

L218 19 DUE RE: L217 L215 L213 L216 L214 (0 DUPLICATES REMOVED)

ANSWERS '1-6' FROM FILE MEDLINE
ANSWERS '7-10' FROM FILE DRUGU
ANSWERS '11-15' FROM FILE CAPLUS
ANSWERS '16-17' FROM FILE WPIX
ANSWERS '18-19' FROM FILE EMBASE

=> d iell 1-10; d ibib abe hit 11-15; d ibib abeq tech hitstr 16-17; d iell 18-19; fil hom

L218 ANSWER 1 OF 19 MEDLINE on STN
ACCESSION NUMBER: 2006205320 MEDLINE Full-text
DOCUMENT NUMBER: PubMed ID: 16612164
TITLE: Efficacy of controlled-release oxycodone for dyspnea in cancer patients--three case series.
AUTHOR: Shinjo Takuji; Okada Masakuni
CORPORATE SOURCE: Dept. of Palliative Care Unit, Shakahoken Kobe Central Hospital.
SOURCE: Onc to kagaku ryoho. Cancer & chemotherapy. (2006 Apr) Vol. 33, No. 4, pp. 529-32.
Journal code: 7810034. ISSN: 0365-0684.
PUB. COUNTRY: Japan
DOCUMENT TYPE: (CASE REPORTS)
Language: English
FILE SEGMENT: Priority Journals
ENTRY MONTH: 200605
ENTRY DATE: Entered STN: 14 Apr 2006
Last Updated on STN: 10 May 2006
Entered Medline: 9 May 2006

ABSTRACT:
Dyspnea is a common symptom in patients with advanced cancer. Systemic morphine administration has been reported as an effective pharmacological treatment to control dyspnea. However, there have been few reports on similar effects of alternative opioids except for morphine. To evaluate the effect of controlled-release oxycodone on the relief of dyspnea, we investigated three cases with opioid substitution from subcutaneous morphine to oral oxycodone. In all cases, both opioids provided equivalent effects for the palliation of cancer dyspnea with no significant adverse effects. Future studies in the appropriate clinical designs will be needed to confirm our findings.

CONTROLLED TERM: Check Tag: Female; Male
Administration, Oral
Aged
*Analgesics, Opioid: AD, administration & dosage
Delayed-Action Preparations
*Dyspnea: DR, drug therapy
Dyspnea: ET, etiology
English Abstract
Humans
Injections, Subcutaneous
Lung Neoplasms: CO, complications
*Lung Neoplasms: PP, physiopathology
Middle Aged
Morphine: AD, administration & dosage
*Oxycodone: AD, administration & dosage
CAS REGISTRY NO.: 57-27-2 (Morphine); 76-42-6 (Oxycodone)
CHEMICAL NAME: O (Analgesics, Opioid); O (Delayed-Action Preparations)

L218 ANSWER 2 OF 19 MEDLINE on STN
ACCESSION NUMBER: 2005170367 MEDLINE Full-text
DOCUMENT NUMBER: PubMed ID: 15801946

TITLE: Co-administration of oxycodone and morphine and analgesic synergy re-examined.
AUTHOR: Smith Maree T; de la Iglesia Felix A'
SOURCE: British journal of clinical pharmacology. (2005 Apr) Vol. 59, No. 4, pp. 486-7; author reply 487-8.
Journal code: 7503323. ISSN: 0306-5251.
Comment: From: Br J Clin Pharmacol. 2004 Sep;58(3):235-42.
PubMed ID: 15327582
England: United Kingdom
Commentary
Letter
Language: English
FILE SEGMENT: Priority Journals
ENTRY MONTH: 200508
ENTRY DATE: Entered STN: 2 Apr 2005
Last Updated on STN: 2 Aug 2005
Entered Medline: 1 Aug 2005
CONTROLLED TERM: *Analgesics, Opioid: AD, administration & dosage
*Cold
Drug Combinations
Drug Synergism
Humans
*Morphine: AD, administration & dosage
*Nociceptors: DE, drug effects
*Oxycodone: AD, administration & dosage
*Pain: PC, prevention & control
57-27-2 (Morphine); 76-42-6 (Oxycodone)
O (Analgesics, Opioid); O (Drug Combinations)

L218 ANSWER 3 OF 19 MEDLINE on STN
ACCESSION NUMBER: 2004422001 MEDLINE Full-text
DOCUMENT NUMBER: PubMed ID: 15327582
TITLE: Can coadministration of oxycodone and morphine produce analgesic synergy in humans? An experimental cold pain study.
AUTHOR: Grach Michael; Massalha Wattan; Pud Dorit; Adler Rivka; Eisenberg Elan
CORPORATE SOURCE: Department of Anesthesiology, Carmel Hospital, Haifa, Israel.
SOURCE: British journal of clinical pharmacology. (2004 Sep) Vol. 58, No. 3, pp. 235-42.
Journal code: 7503323. ISSN: 0306-5251.
Comment: In: Br J Clin Pharmacol. 2005 Apr;59(4):486-7; author reply 487-8. PubMed ID: 15801946
PUB. COUNTRY: England: United Kingdom
DOCUMENT TYPE: (CLINICAL TRIAL)
Language: English
FILE SEGMENT: Priority Journals
ENTRY MONTH: 200412
ENTRY DATE: Entered STN: 26 Aug 2004
Last Updated on STN: 20 Dec 2004
Entered Medline: 17 Dec 2004

ABSTRACT:
AIMS: The coadministration of subantinociceptive doses of oxycodone with morphine has recently been shown to result in a synergistic antinociceptive effect in rats. The present study was aimed to investigate the possibility that coadministration of morphine and oxycodone can produce a similar synergistic effect in humans exposed to an experimental model

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of cold pressor test (CPT). METHODS: The enriched enrollment design was used to exclude 'atonic' and 'placebo responders' in a single-blind fashion. 'Monstoxic', placebo 'nonresponder' female volunteers ($n = 30$) were randomly assigned to receive 0.5 mg kg⁻¹ oral morphine sulphate, 0.5 mg kg⁻¹ oral oxycodone hydrochloride, and the combination of 0.25 mg kg⁻¹ morphine sulphate with 0.25 mg kg⁻¹ oxycodone hydrochloride, 1 week apart from each other, in a double-blind crossover design. Latency to pain onset (threshold), pain intensity (VAS), and pain tolerance (time until removal of the hand from the water) were measured six times over a 3-h period, subsequent to the administration of each medication, and were used to assess their antinociceptive effect. RESULTS: The combination produced a significantly higher effect on latency to pain onset than that of morphine alone (difference in mean postbaseline value 2.1; 95% confidence interval (CI) 0.48, 3.9; $P < 0.01$) but the effect was nonsignificantly smaller than that of oxycodone alone. Similarly, the effect of the combination on pain tolerance was significantly larger than that of morphine alone (combination difference 8.4; 95% CI 2.5, 14.3; $P = 0.007$), whereas oxycodone alone caused a nonsignificantly larger effect than that of the combination treatment. Comparisons of pain magnitude failed to show any significant differences between the three treatments. CONCLUSIONS: These results indicate that at the doses tested, morphine and oxycodone do not produce synergistic antinociceptive effects in healthy humans exposed to the CPT.

CONTROLLED TERM: Check Tags: Female

- Adolescent
- Adult
- *Analgesics, Opioid: AD, administration & dosage
- *Cold
- Cross-Over Studies
- Double-Blind Method
- Drug Combinations
- Drug Synergism
- Humans
- *Morphine: AD, administration & dosage
- *Oxycodone: AD, administration & dosage
- *Pain: PC, prevention & control

CAS REGISTRY NO.: 57-27-2 (Morphine); 76-42-6 (Oxycodone)
CHEMICAL NAME: 0 (Analgesics, Opioid); 0 (Drug Combinations)

L218 ANSWER 4 OF 19 MEDLINE on STN

ACCESSION NUMBER: 20031606935 MEDLINE Full-text

DOCUMENT NUMBER: PubMed ID: 14557380

TITLE: Opioid interactions in rhesus monkeys: effects of delta + mu and delta + kappa agonists on schedule-controlled responding and thermal nociception.

AUTHOR: Stevenson Glenn W; Foltz John B; Linsenmayer David C; Rice Kenner C; Negus S Stevens

CORPORATE SOURCE: Alcohol and Drug Abuse Research Center, Harvard Medical School, McLean Hospital, 115 Mill St., Belmont, MA 02478-9106, USA.

CONTRACT NUMBER: P01-DA14528 (NIDA)
R01-DA11460 (NIDA)
T32-DA07252 (NIDA)

SOURCE: The Journal of pharmacology and experimental therapeutics, (2003 Dec) Vol. 307, No. 3, pp. 1054-64. Electronic

Publication: 2003-10-13.

Journal code: 0376362. ISSN: 0022-3565.

PUB. COUNTRY: United States

DOCUMENT TYPE: Journal; Article; (JOURNAL ARTICLE)

LANGUAGE: English

10/661458

FILE SEGMENT: Priority Journals
ENTRY MONTH: 200401
ENTRY DATE: Entered STN: 24 Dec 2003
Last Updated on STN: 30 Jan 2004
Entered Medline: 29 Jan 2004

ABSTRACT:

Agonists at delta, mu, and kappa opioid receptors produce interacting effects in rodents and nonhuman primates. To further evaluate the determinants of these interactions, this study examined the effects of mixtures of delta + mu and delta + kappa agonists in rhesus monkeys ($n = 4-5$) using two behavioral procedures, an assay of schedule-controlled responding for food reinforcement and an assay of thermal nociception. Results were analyzed using dose-addition analysis. In the assay of schedule-controlled responding, the delta agonist (+)-4-[(alphaR)-alpha-((2S,5R)-4-allyl-2,5-dimethyl-1-piperazinyl)-3-methoxybenzyl]-N,N-diethyl-benzamide (SNC80); the mu agonists methadone, fentanyl, morphine, and nalbuphine; and the kappa agonists (Salpa, 7alpha, 7beta, 11-methyl-N-(7-(1-pyrrolidinyl)-1-oxaspiro[4.5]dec-8-yl)benzenacetamide (US, 593) and bremazocine all dose-dependently decreased rates of food-maintained responding when administered alone. Fixed ratio mixtures of SNC80 + mu agonists produced additive or subadditive effects, whereas SNC80 + kappa agonist mixtures produced only additive effects. In the assay of thermal nociception, SNC80 produced no measurable effects when administered alone, whereas mu and kappa agonists produced dose-dependent antinociception. SNC80 + mu agonist mixtures produced superadditive effects manifested as leftward shifts in mu agonist dose-effect curves. This synergism was antagonized by the delta-selective antagonist naltrindole, suggesting that SNC80-induced enhancement of mu agonist antinociception was delta receptor-mediated. SNC80 did not enhance the antinociceptive effects of the highly selective kappa agonist US, 593, and it produced only a marginal enhancement of antinociception produced by the less selective kappa agonist bremazocine. These results suggest that delta agonists may selectively enhance the antinociceptive effects of mu agonists in rhesus monkeys. These results also confirm that opioid agonist interactions may depend on the receptor selectivity and relative doses of the agonists and on the experimental endpoint.

CONTROLLED TERM: Check Tags: Male

- *Analgesics, Opioid: PD, pharmacology
- Animals
- Benzamides: PD, pharmacology
- Benzeneacetamides: PD, pharmacology
- Benzomorphans: PD, pharmacology
- *Conditioning, Operant: DE, drug effects
- Dose-Response Relationship, Drug
- Drug Interactions

Heat

Macaca mulatta

*Naltrrexone: AA, analogs & derivatives

Naltrrexone: PD, pharmacology

Narcotic Antagonists: PD, pharmacology

*Pain: PX, psychology

Piperazines: PD, pharmacology

Pyrrolidinines: PD, pharmacology

*Receptors, Opioid, delta: AG, agonists

*Receptors, Opioid, kappa: AG, agonists

*Receptors, Opioid, mu: AG, agonists

Reinforcement Schedule

Research Support, U.S. Gov't, P.H.S.

CAS REGISTRY NO.: 111555-53-4 (Naltrindole); 156727-74-1 (4-(alpha-(4-allyl-2,5-dimethyl-1-piperazinyl)-3-methoxybenzyl)-N,N-diethylbenzamide); 16580-41-3 (Naltrrexone); 75684-07-0 (Bremazocine); 96744-75-1 (U 69593)

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CHEMICAL NAME: 0 (Analgesics, Opioid); 0 (Benzamides); 0 (Benzeneacetamides); 0 (Benzomorphans); 0 (Narcotic Antagonists); 0 (Piperazines); 0 (Pyrrolidinines); 0 (Receptors, Opioid, delta); 0 (Receptors, Opioid, kappa); 0 (Receptors, Opioid, mu)

L218 ANSWER 5 OF 19 MEDLINE on STN

ACCESSION NUMBER: 2000107229 MEDLINE Full-text

DOCUMENT NUMBER: PubMed ID: 10640321

TITLE: The antitussive activity of delta-opioid receptor stimulation in guinea pig.

AUTHOR: Kotzer C J; Hay D W; Dondio G; Giardina G; Petrillo P; Underwood D C

CORPORATE SOURCE: Department of Pulmonary Pharmacology, SmithKline Beecham Pharmaceuticals, King of Prussia, Pennsylvania, USA.

SOURCE: The Journal of pharmacology and experimental therapeutics, (2000 Feb) Vol. 292, No. 2, pp. 803-9.

Journal code: 0376362. ISSN: 0022-3565.

PUB. COUNTRY: United States

DOCUMENT TYPE: Journal; Article; (JOURNAL ARTICLE)

LANGUAGE: English

FILE SEGMENT: Priority Journals

ENTRY MONTH: 200002

ENTRY DATE: Entered STN: 9 Mar 2000

Last Updated on STN: 9 Mar 2000

Entered Medline: 22 Feb 2000

ABSTRACT: In this study, the activity of the delta-opioid receptor subtype-selective agonist, SB 227122, was investigated in a guinea pig model of citric acid-induced cough. Parenteral administration of selective agonists of the delta-opioid receptor (SB 227122), mu-opioid receptor (codeine and hydrocodone), and kappa-opioid receptor (BRL 52974) produced dose-related inhibition of citric acid-induced cough with ED₅₀ values of 7.3, 5.2, 5.1, and 5.3 mg/kg, respectively. The nonselective opioid receptor antagonist, naloxone (3 mg/kg, i.m.), attenuated the antitussive effects of codeine or SB 227122, indicating that the antitussive activity of both compounds is opioid receptor-mediated. The delta-receptor antagonist, SB 244525 (10 mg/kg, i.p.), inhibited the antitussive effect of SB 227122 (20 mg/kg, i.p.). In contrast, combined pretreatment with beta-funaltrexamine (mu-receptor antagonist; 20 mg/kg, s.c.) and norbinaltorphimine (kappa-receptor antagonist; 20 mg/kg, s.c.), at doses that inhibited the antitussive activity of mu- and kappa-receptor agonists, respectively, was without effect on the antitussive response of SB 227122 (20 mg/kg, i.p.). The sigma-receptor antagonist, rimecazole (3 mg/kg, i.p.) inhibited the antitussive effect of dextromethorphan (30 mg/kg, i.p.), a sigma-receptor agonist, but not that of SB 227122. These studies provide compelling evidence that the antitussive effects of SB 227122 in this guinea pig cough model are mediated by agonist activity at the delta-opioid receptor.

CONTROLLED TERM: Check Tags: Male

- Animals
- CHO Cells
- Carbazoles: PD, pharmacology
- Cell Lines
- Cloning, Organism
- Codine: PD, pharmacology
- *Cough: PC, prevention & control
- Cricetines
- Dextromethorphan: PD, pharmacology
- Disease Models, Animal
- Dose-Response Relationship, Drug

10/661458

Drug Interactions

Guinea Pigs

Humans

Hydrocodone: PD, pharmacology

*Levallophan: AA, analogs & derivatives

Levallophan: TU, therapeutic use

Naloxone: PD, pharmacology

*Narcotic Antagonists: PD, pharmacology

Protein Binding

Pyridines: PD, pharmacology

*Pyrolo: TU, therapeutic use

Pyrrolidinines: PD, pharmacology

*Receptors, Opioid, delta: AG, agonists

*Receptors, Opioid, delta: DE, drug effects

*Receptors, Opioid, delta: PH, physiology

Receptors, Opioid, kappa: AG, agonists

Receptors, Opioid, kappa: DE, drug effects

Receptors, Opioid, kappa: PH, physiology

Receptors, opioid, mu: AG, agonists

Receptors, Opioid, mu: DE, drug effects

Receptors, Opioid, mu: PH, physiology

125-29- (Hydrocodone); 125-71-3 (Dextromethorphan);

145544-79-2 (BRL 52974); 152-02-3 (Levallophan); 465-65-6 (Naloxone); 75859-04-0 (Rimecazole); 76-57-3 (Codeine)

(Carbazole); 0 (Narcotic Antagonists); 0 (Pyridines); 0 (Pyrroles); 0 (Pyrrolidinines); 0 (Receptors, Opioid, kappa); 0 (Receptors, Opioid, mu); 0 (SB 227122)

L218 ANSWER 6 OF 19 MEDLINE on STN

ACCESSION NUMBER: 2000133083 MEDLINE Full-text

DOCUMENT NUMBER: PubMed ID: 10666549

TITLE: Co-administration of sub-antinociceptive doses of oxycodone and morphine produces marked antinociceptive synergy with reduced CNS side-effects in rats.

AUTHOR: Rose F B; Wallis S C; Smith M T

CORPORATE SOURCE: School of Pharmacy, The University of Queensland, St Lucia, Brisbane, Australia.

SOURCE: Pain, (2000 Feb) Vol. 84, No. 2-3, pp. 421-8.

JOURNAL CODE: 750866. ISSN: 0304-3959.

PUB. COUNTRY: Netherlands

DOCUMENT TYPE: Journal; Article; (JOURNAL ARTICLE)

LANGUAGE: English

FILE SEGMENT: Priority Journals

ENTRY MONTH: 200003

ENTRY DATE: Entered STN: 27 Mar 2000

Last Updated on STN: 27 Mar 2000

Entered Medline: 16 Mar 2000

ABSTRACT:

Oxycodone and morphine are structurally related, strong opioid analgesics, commonly used to treat moderate to severe pain in humans. Although it is well-established that morphine is a mu-opioid agonist, this is not the case for oxycodone. Instead, our recent studies have shown that oxycodone appears to be a kappa-opioid agonist (Rose and Smith, 1997). In the current study, we now show that co-administration of sub-antinociceptive doses of oxycodone (putative kappa-opioid agonist) with morphine (mu-opioid agonist) to rats by both the intracerebroventricular and by systemic routes (intraperitoneal and subcutaneous), results in markedly increased (synergistic) levels of antinociception. Behaviourally, rats co-administered sub-antinociceptive doses of oxycodone and morphine were similar to control rats dosed with saline.

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whereas rats that received equi-potent doses of either opioid alone, were markedly sedated. These results suggest that co-administration of sub-analgesic doses of oxycodone and morphine to patients may provide excellent pain relief with a reduction in opioid-related CNS side-effects. Controlled clinical trials in appropriate patient populations are required to evaluate this possibility.(1)

CONTROLLED TERM:

Check Tags: Male
 Analgesics, Opioid: AD, administration & dosage
 *Analgesics, Opioid: PD, pharmacology
 Animals
 Behavior, Animal: DE, drug effects
 Central Nervous System: DE, drug effects
 Dose-Response Relationship, Drug
 Drug Combinations
 Drug Synergism
 Injections, Intraperitoneal
 Injections, Intraventricular
 Injections, Subcutaneous
 Morphine: AD, administration & dosage
 *Morphine: PD, pharmacology
 *Nociceptors: DE, drug effects
 Oxycodeone: AD, administration & dosage
 *Oxycodeone: PD, pharmacology
 Rats
 Rats, Sprague-Dawley
 Research Support, Non-U.S. Gov't
 CAS REGISTRY NO.: 57-27-2 (Morphine); 76-42-6 (Oxycodeone)

CHEMICAL NAME:
 0 (Analgesics, Opioid); 0 (Drug Combinations)

L218 ANSWER 7 OF 19 DRUGU COPYRIGHT 2006 THE THOMSON CORP on STN

ACCESSION NUMBER: 1995-21708 DRUGU T Full-text
 TITLE: A pain syndrome associated with large adrenal masses in patients with lung cancer.

AUTHOR: Berger M S; Cooley M E; Abraham J L

CORPORATE SOURCE: Univ. Pennsylvania

LOCATION: Philadelphia, Pa., USA

SOURCE: J.Pain Symptom Manage. (10, No. 2, 161-66, 1995) 2 Fig. 13

Ref.

CODEN: JPSM2U ISSN: 0885-3924
 AVAIL. OF DOC.: Hematology-Oncology Division, Philadelphia VA Medical Center, University and Woodlands Avenues, Philadelphia, PA, U.S.A.

LANGUAGE: English

DOCUMENT TYPE: Journal

ABSTRACT:

Case histories are reported of 2 patients with lung cancer who had a pain syndrome caused by large adrenal metastases. Patient 1 had a poor response to radiation, controlled-release p.o. morphine and acetaminophen-oxycodone. He responded to chemotherapy with cyclophosphamide (Cytosan), Adriamycin and vincristine (CAV). He was given hydrocortisone for orthostatic hypotension. Hip pain developed and he died. Patient 2 was treated with controlled-release p.o. morphine but pain progressed and he died. 23 Previously recorded cases were reviewed.

SECTION HEADING: T Therapeutics

CLASSIF. CODE: 43 Analgesics, NSAIDs
 44 Narcotics

51 Chemotherapy - clinical

CONTROLLED TERM:

ADRENAL *TR; METASTASIS *TR; ADRENOPATHY *TR; NEOPLASM *TR; PAIN *TR; LUNG *OC; SMALL-CELL *OC; LARGE-CELL *OC; NEOPLASM *OC; HYDROCORTISONE *RC; CASE-HISTORY *FT; IN-VIVO *FT; RADIOTHERAPY *FT; CONCOMITANT-DISEASE *FT; EXITUS *FT; CASES *FT

(01) MORPHINE *TR; MORPHINE *RN; ANALGESIC *FT; DRUG *FT; P.O. *FT; PHARM.PREP. *FT; ANALGESICS *FT; NARCOTICS *FT; SEDATIVES *FT; 57-27-2 *FT; TR *FT

CAS REGISTRY NO.: 57-27-2
 (02) OXYCODEONE *TR; OXYCODEONE *RN; COMB. PREP. *FT; P.O. *FT; ANALGESIC *FT; ANALGESICS *FT; NARCOTICS *FT; SEDATIVES *FT; 76-42-6 *FT; TR *FT

CAS REGISTRY NO.: 76-42-6
 (03) PARACETAMOL *TR; PARACETAMOL *RN; COMB.PREP. *FT; ANALGESIC *FT; P.O. *FT; ANALGESICS *FT; ANTIPIRETICS *FT; 103-90-2 *FT; TR *FT

CAS REGISTRY NO.: 103-90-2
 (04) CYCLOPHOSPHAMIDE *TR; CYTOXAN *TR; CYCLOPHOS *RN; CYTOSTATIC *FT; CYTOSTATIC-COMB. *FT; COMB. *FT; CYTOSTATICS *FT; IMMUNOSUPPRESSIVES *FT; 50-18-0 *FT; TR *FT

CAS REGISTRY NO.: 50-18-0
 (05) DOXORUBICIN *TR; ADRIAMYCIN *TR; DOXORUBICIN *RN; CYTOSTATIC *FT; CYTOSTATIC-COMB. *FT; COMB. *FT; ANTIOTICOS *FT; CYTOSTATICS *FT; 23214-92-8 *FT; TR *FT

CAS REGISTRY NO.: 23214-92-8
 (06) VINCRISTINE *TR; VINCRISTI *RN; CYTOSTATIC *FT; CYTOSTATIC-COMB. *FT; COMB. *FT; CYTOSTATICS *FT; 57-22-7 *FT; TR *FT

CAS REGISTRY NO.: 57-22-7
 FIELD AVAIL.: AB: LA; CT
 FILE SEGMENT: Literature

L218 ANSWER 8 OF 19 DRUGU COPYRIGHT 2006 THE THOMSON CORP on STN

ACCESSION NUMBER: 1993-55507 DRUGU T S Full-text
 TITLE: A Risk-Benefit Appraisal of Injectable NSAIDs in the Management of Postoperative Pain.

AUTHOR: Huutinen L S; Laitinen J O; Salomaki T E

LOCATION: Kuopio, Oulu, Finland

SOURCE: Drug Safety (9, No. 5, 380-93, 1993) 3 Tab. 124 Ref.

ISSN: 0114-5916
 AVAIL. OF DOC.: Department of Anaesthesiology, University Hospital, P.O.B. 1777, SF-70211 Kuopio, Finland.

LANGUAGE: English
 DOCUMENT TYPE: Journal

ABSTRACT:

Injectable NSAIDs in the management of postoperative pain are reviewed, with reference to their mode of action, the use of indometacin (IN), diclofenac (DI), ketorolac (KE) and other NSAIDs for acute pain, the adverse effects of NSAIDs on the GI system, coagulation and renal and other adverse effects. Somnolence, dry mouth and GI effects are the commonest adverse events with KE. Interactions occur between NSAIDs and anticoagulants, diuretics, beta-blockers and lithium. Parenteral NSAIDs, particularly IN, DI and KE, have a clear role in the management of postoperative pain. Their efficacy is well proved in orthopedic surgery. Their use is contraindicated in patients with a history of

asthma , allergy, renal pathology or peptic ulceration.

SECTION HEADING: T Therapeutics
 S Adverse Effects

CLASSIF. CODE: 35 Adverse Reactions
 43 Analgesics, NSAIDs
 44 Narcotics
 66 Drug Interactions
 69 Reviews

CONTROLLED TERM:
 PAIN *TR; POSTOPERATIVE *TR; IN-VIVO *FT; CASES *FT; INJECTION *FT; ANTIINFLAMMATORY *FT; REVIEW *FT; RISK-FACTOR *FT
 (01) ANTIINFLAMMATORIES *FT; MAIN-TOPIC *FT; TR *FT; AE *FT; DI
 (02) INDOMESTACIN *TR; INDOMESTACIN *AE; DICLOFENAC *AE; DICLOFENAC *TR; KETOROLAC *TR; KETOROLAC *AE; INDOMESTACIN *DI; DICLOFENAC *DI; KETOROLAC *DI; OXYCODEONE *TR; PENTHADINE *TR; PENTHADINE *TR; MORPHINE *TR; MORPHINE *AE; ASPIRIN-LYSINE-SALT *TR; KETOPROFEN *TR; INDOPOFEN *TR; TERNOXICAM *TR; PIROXICAM *AE; OXYCODEONE *AE; PENTHADINE *AE; ASPIRIN *AE; ALFENTANIL *AE; MODE-OF-ACT. *FT; ORTHOPEDICS *FT; SURGERY *FT; CONTRAINDICATION *FT; DRUG-COMPARISON *FT; I.V. *FT; INJECTION *FT; TR *FT; AE *FT; DI *FT
 FIELD AVAIL.: AB: LA; CT
 FILE SEGMENT: Literature

L218 ANSWER 9 OF 19 DRUGU COPYRIGHT 2006 THE THOMSON CORP on STN

ACCESSION NUMBER: 1991-28479 DRUGU S Full-text
 TITLE: A Prospective Study of Hospital Admissions Due to Drug Reactions.

AUTHOR: Larmour I; Dolphin R G; Baxter H; Morrison S; Hooke D H; McGrath B P

LOCATION: Melbourne, Australia
 SOURCE: Aust.J.Hosp.Pharm. (21, No. 2, 90-95, 1991) 2 Fig. 4 Tab. 14

Ref.

CODEN: AUPHAI ISSN: 0310-6810
 AVAIL. OF DOC.: Manager of Pharmaceutical Services, Monash Medical Center, Prince Henry's Hospital, St. Kilda Road, Melbourne, Vic. 3004, Australia.

LANGUAGE: English
 DOCUMENT TYPE: Journal

ABSTRACT:

67 Drugs were implicated in adverse drug reactions (ADRs) in 136/5623 hospital admissions (2.4%) in a 6-mth prospective study. Drugs included piroxicam, diclofenac, indometacin, diflunisal, ketoprofen, naproxen, cimetidine, doxycycline, warfarin, aspirin, dipyradomole, hydralazine, cyclopentiazide, atenolol, metoprolol, digoxin, amlodarone, verapamil, nifedipine, chlorothiazide, methylthiazide, theophylline, allopurinol, ranitidine, methotrexate, glibenclamide, metformin, prochlorperazine, oxycodone, bromocriptine, thioridazine, naproxen, bleomycin, promethazine, morphine, allopurinol, co-trimoxazole, (trimethoprim + sulfamethoxazole), cyclophosphamide. Most ADRs were GI bleeding and cardiovascular complications;

5 were fatal.

SECTION HEADING: S Adverse Effects

CLASSIF. CODE: 18 Hematological
 35 Adverse Reactions
 43 Analgesics, NSAIDs
 58 Vasoactive
 66 Drug Interactions

CONTROLLED TERM:

IN-VIVO *FT; CASES *FT; COMB. *FT
 (01) CYCLOPHOSPHAMIDE *DI; CYCLOPHOSPHAMIDE *AE; MARROW-DISEASE *AE; HEMOPTYSIS *AE; HEMORRHAGE *AE; CYSTITIS *AE; BLADDER-DISEASE *AE; WARFARIN *DI; ASPIRIN *DI; CYTOSTATICS *FT; IMMUNOSUPPRESSIVES *FT; CYCLOPHOSPHAMIDE *RN; AE *FT; DI *FT
 CAS REGISTRY NO.: 50-18-0
 (02) WARFARIN *DI; WARFARIN *AE; PANCYTOPENIA *AE; MARROW-DISEASE *AE; HEMOPTYSIS *AR; MELENA *AE; HEMORRHAGE *AR; CYSTITIS *AE; HEMORRHAGE *AS; ANEMIA *AB; GASTROENTEROPATHY *AB; HEMORRHAGE *AB; EMESES *AB; GASTROENTEROPATHY *AB; DEHYDRATION *DI; ANEMIA *AB; BLADDER-DISEASE *AB; TRIMETHOPRIM *DI; DOXYCYCLINE *DI; SULFAMETHOXAZOLE *DI; ASPIRIN *DI; PREDNISOLONE *DI; Cimetidine *DI; CYCLOPHOSPHAMIDE *DI; RODENTICIDES *FT; ANTI COAGULANTS *FT; WARFARIN *RN; DI *FT; AE *FT
 CAS REGISTRY NO.: 5543-58-8
 (03) ASPIRIN *DI; ASPIRIN *AE; PANCYTOPENIA *AE; MARROW-DISEASE *AB; HEMOPTYSIS *AB; HEMORRHAGE *AB; CYSTITIS *AE; HEMATOMESESIS *AB; MELENA *AB; GASTROENTEROPATHY *AB; HEMORRHAGE *AS; ANEMIA *AB; BLADDER-DISEASE *AB; DICLOFENAC *DI; DIPYRIDAMOLE *DI; INDOMETACIN *DI; KETOPROFEN *DI; NAPROXEN *DI; PIROXICAM *DI; WARFARIN *DI; PREDNISOLONE *DI; CYCLOPHOSPHAMIDE *DI; ANALGESICS *FT; ANTI PYRETICS *FT; ANTI RHEUMATICS *FT; ANTI AGGREANTS *FT; PROSTAGLANDIN-ANTAGONISTS *FT; ASPIRIN *RN; DI *FT; AE *FT
 CAS REGISTRY NO.: 5543-58-8
 (04) TRIMETHOPRIM *DI; TRIMETHOPRIM *AB; HEMATOMESESIS *AB; ANEMIA *AB; CEREBROVASCULAR-DISEASE *AB; MELENA *AB; GASTROENTEROPATHY *AB; HEMORRHAGE *AB; DEHYDRATION *AB; MUCOSITIS *AB; WARFARIN *DI; Cimetidine *DI; DIPYRIDAMOLE *DI; INDOMETACIN *DI; KETOPROFEN *DI; NAPROXEN *DI; PIROXICAM *DI; WARFARIN *DI; ASPIRIN *DI; PREDNISOLONE *DI; CYCLOPHOSPHAMIDE *DI; CIMETIDINE *DI; PREDNISOLONE *DI; CYCLOPHOSPHAMIDE *DI; METOTREXATE *DI; COMB. PREP. *FT; TRIMETHOPRIM *RN; DI *FT; AE *FT
 CAS REGISTRY NO.: 738-76-2
 (05) SULFAMETHOXAZOLE *AB; HEMATOMESESIS *AB; ANEMIA *AB; CEREBROVASCULAR-DISEASE *AB; MELENA *AB; GASTROENTEROPATHY *AB; DEHYDRATION *AB; MUCOSITIS *AB; WARFARIN *DI; Cimetidine *DI; DOXYCYCLINE *DI; ASPIRIN *DI; PREDNISOLONE *DI; CYCLOPHOSPHAMIDE *DI; METOTREXATE *DI; COMB. PREP. *FT; ANTI SEPTICS *FT; SULFAMOXIA *RN; AE *FT
 CAS REGISTRY NO.: 723-46-6
 (06) DICLOFENAC *AB; DICLOFENAC *DI; HEMATOMESESIS *AB; MELENA *AB; GASTROENTEROPATHY *AB; HEMORRHAGE *AB; ANEMIA *AB; HEMOPTYSIS *AB; DICLOFENAC *AB; DICLOFENAC *DI; ANTI INFLAMMATORIES *FT; ANALGESICS *FT; PROSTAGLANDIN-ANTAGONISTS *FT; DICLOFENAC *RN; AE *FT

CAS REGISTRY NO.: 15307-86-5
BLEOMYCIN *AE; PULMONARY-FIBROSIS *AE;
PNEUMOPATHY *AE; NEUTROPENIA *AE; MARROW-DISEASE *AE;
THROMBOCYTOPENIA *AE; ANTIOTIOTICS *PT; CYTOSTATICS *PT;
BLEOMYCIN *RN; AE *PT

CAS REGISTRY NO.: 11056-06-7
[68] NAPROXEN *AE; HEMATEMESIS *AE; MELENA *AE; GASTROENTEROPATHY
*AE; HEMORRHAGE *AE; PROSTAGLANDIN-ANTAGONISTS *PT;
ANTIINFLAMMATORIES *PT; ANALGESICS *PT; ANTIPIRETICS *PT;
NAPROXEN *RN; AE *PT

CAS REGISTRY NO.: 12204-53-1
[69] DIGOXIN *DI; DIGOXIN *AE; BRADYCARDIA *AE; ANOREXIA *AE;
HEART-BLOCK *AE; ARRHYTHMIA *AE; CARDIOPATHY *AE; AMIODARONE
*DI; VERAPAMIL *DI; ATENOLOL *DI; NIFEDIPINE *DI;
CARDIOLYCOSES *PT; CARDIANTS *PT; DIGOXIN *RN; DI *PT; AE
*PT

CAS REGISTRY NO.: 20830-75-5
[70] Cimetidine *AE; Cimetidine *DI; HEMATEMESIS *AE; ANEMIA *AE;
CEREBROVASCULAR-DISEASE *AE; WARFARIN *DI; TRIMETHOPRIM *DI;
SULFAMETHOXAZOLE *DI; ANTIHISTAMINES-H2 *PT; ANTIULCERS *PT;
GASTRIC-SECRETION-INHIBITORS *PT; Cimetidine *RN; AE *PT; DI
*PT

CAS REGISTRY NO.: 51481-61-9
[71] AMIODARONE *DI; AMIODARONE *AE; BRADYCARDIA *AE; ARRHYTHMIA
*AE; CARDIOPATHY *AE; ANOREXIA *AE; DIGOXIN *DI; VERAPAMIL
*DI; CALCIUM-ANTAGONISTS *PT; CARDIANTS *PT; ANTIARRHYTHMICS
*PT; AMIODARONE *RN; DI *PT; AE *PT

CAS REGISTRY NO.: 1851-25-3
[72] VERAPAMIL *DI; VERAPAMIL *AE; ARRHYTHMIA *AE; CARDIOPATHY
*AE; ANOREXIA *AE; DIGOXIN *DI; AMIODARONE *DI; CARDIANTS
*PT; CALCIUM-ANTAGONISTS *PT; VERAPAMIL *RN; DI *PT; AE *PT

CAS REGISTRY NO.: 52-53-9
[73] ATENOLOL *DI; ATENOLOL *AE; HEART-BLOCK *AE; ARRHYTHMIA *AE;
CARDIOPATHY *AE; DIGOXIN *DI; NIFEDIPINE *DI;
SYMPATHOLYTICS-BETA *PT; HYPOTENSIVES *PT; ATENOLOL *RN; DI
*PT; AE *PT

CAS REGISTRY NO.: 29122-68-7
[74] NIFEDIPINE *DI; NIFEDIPINE *AE; BRADYCARDIA *AE; ARRHYTHMIA
*AE; CARDIOPATHY *AE; DIGOXIN *DI; ATENOLOL *DI; CARDIANTS
*PT; CALCIUM-ANTAGONISTS *PT; NIFEDIPINE *RN; DI *PT; AE *PT

CAS REGISTRY NO.: 21829-35-4
[75] METHYLDOPA *DI; METHYLDOPA *AE; ORTHOSTATIC *AE; HYPOTENSION
*AE; VASCULAR-DISEASE *AE; PERIPHERAL-NERVE-DISEASE *AE;
CHLOROTHIAZIDE *DI; VERAPAMIL *DI; CHLORPROMAZINE *DI;
HYDRALAZINE *DI; HYPOTENSIVES *PT; SYMPATHOMIMETICS-ALPHA
*PT; METHYLDOPA *RN; DI *PT; AE *PT

CAS REGISTRY NO.: 556-30-6
[76] CHLOROTHIAZIDE *DI; CHLOROTHIAZIDE *AE; ORTHOSTATIC *AE;
HYPOTENSION *AE; VASCULAR-DISEASE *AE; PERIPHERAL-NERVE-
DISEASE *AE; METHYLDOPA *DI; VERAPAMIL *DI; CHLORPROMAZINE
*DI; CARBONIC-ANHYDRASE-INHIBITORS *PT; DIURETICS *PT;
HYPOTENSIVES *PT; CHLOROTHIAZIDE *RN; DI *PT; AE *PT

CAS REGISTRY NO.: 58-94-6
[77] CHLORPROMAZINE *DI; CHLORPROMAZINE *AE; ORTHOSTATIC *AE;
HYPOTENSION *AE; VASCULAR-DISEASE *AE; PERIPHERAL-NERVE-
DISEASE *AE; METHYLDOPA *DI; CHLOROTHIAZIDE *DI; VERAPAMIL
*DI; HYDRALAZINE *DI; PSYCHODERATIVES *PT; NEUROLEPTICS *PT;
SEDATIVES *PT; DOPAMINE-ANTAGONISTS *PT; CALMODULIN-
ANTAGONISTS *PT; CHLORPROZ *RN; DI *PT; AE *PT

CAS REGISTRY NO.: 50-53-3

[18] HYDRALAZINE *DI; HYDRALAZINE *AE; ORTHOSTATIC *AE;
HYPOTENSION *AE; VASCULAR-DISEASE *AE; PERIPHERAL-NERVE-
DISEASE *AE; METHYLDOPA *DI; VERAPAMIL *DI; CHLORPROMAZINE
*DI; METOPROLOL *DI; CYCLOPENTHIAZIDE *DI; HYPOTENSIVES *PT;
HYDRALAZINE *RN; DI *PT; AE *PT

CAS REGISTRY NO.: 86-54-4
[19] CYCLOPENTHIAZIDE *DI; CYCLOPENTHIAZIDE *AE; ORTHOSTATIC *AE;
HYPOTENSION *AE; VASCULAR-DISEASE *AE; PERIPHERAL-NERVE-
DISEASE *AE; METOPROLOL *DI; HYDRALAZINE *DI; DIURETICS *PT;
HYPOTENSIVES *PT; CYCLOPENTHIAZIDE *RN; DI *PT; AE *PT

CAS REGISTRY NO.: 742-20-1
[20] DIFLUNISAL *DI; DIFLUNISAL *AE; HEMATEMESIS *AE; MELENA *AE;
GASTROENTEROPATHY *AE; HEMORRHAGE *AE; ANEMIA *AE; PIROXICAM
*DI; ANALGESICS *PT; ANTIINFLAMMATORIES *PT; ANTIPIRETICS
*PT; PROSTAGLANDIN-ANTAGONISTS *PT; DIFLUNISAL *RN; DI *PT; AE
*PT

CAS REGISTRY NO.: 22494-42-4
[21] PIROXICAM *DI; PIROXICAM *AE; HEMATEMESIS *AE; MELENA *AE;
GASTROENTEROPATHY *AE; HEMORRHAGE *AE; ANEMIA *AE; DIFLUNISAL
*DI; ANTIINFLAMMATORIES *PT; PROSTAGLANDIN-ANTAGONISTS *PT;
PIROXICAM *RN; DI *PT; AE *PT

CAS REGISTRY NO.: 34322-96-4
[22] METOPROLOL *DI; METOPROLOL *AE; ORTHOSTATIC *AE; HYPOTENSION
*AE; VASCULAR-DISEASE *AE; PERIPHERAL-NERVE-DISEASE *AE;
CYCLOPENTHIAZIDE *DI; HYDRALAZINE *DI; SYMPATHOLYTICS-BETA
*PT; METOPROLOL *RN; DI *PT; AE *PT

CAS REGISTRY NO.: 37350-58-6
[23] DIPYRIDAMOLE *DI; DIPYRIDAMOLE *AE; MELENA *AE;
GASTROENTEROPATHY *AE; HEMORRHAGE *AE; ANEMIA *AE; ASPIRIN
*DI; CARDIANTS *PT; CALCIUM-ANTAGONISTS *PT; ANTIAGGREGANTS
*PT; PROSPHODIESTERASE-INHIBITORS *PT; DIPYRIDAMOLE *RN; DI
*PT; AE *PT

CAS REGISTRY NO.: 50-32-2
[24] INDOMETACIN *DI; INDOMETACIN *AE; MELENA *AE;
GASTROENTEROPATHY *AE; HEMORRHAGE *AE; ASPIRIN *DI;
ANTIINFLAMMATORIES *PT; ANTIPIRETIC *PT; ANTIINFLAMMATORIES
*PT; PROSTAGLANDIN-ANTAGONISTS *PT; INDOMETAC *RN; DI *PT; AE
*PT

CAS REGISTRY NO.: 53-86-1
[25] KETOPROFEN *DI; KETOPROFEN *AE; MELENA *AE; GASTROENTEROPATHY
*AE; HEMORRHAGE *AE; ANEMIA *AE; ASPIRIN *DI;
ANTIINFLAMMATORIES *PT; ANALGESICS *PT; PROSTAGLANDIN-
ANTAGONISTS *PT; KETOPROFEN *RN; DI *PT; AE *PT

CAS REGISTRY NO.: 22071-15-4
[26] DOXYCYCLINE *DI; DOXYCYCLINE *AE; HEMATEMESIS *AE; MELENA
*AE; GASTROENTEROPATHY *AE; HEMORRHAGE *AE; EMESIS *AE;
GASTROENTEROPATHY *AE; DEHYDRATION *AE; WARFARIN *DI;
TRIMETHOPRIM *DI; SULFAMETHOXAZOLE *DI; ANTIBIOTICS *PT;
DOXYCYCLINE *RN; DI *PT; AE *PT

CAS REGISTRY NO.: 56-25-0
[27] PREDNISOLONE *DI; PREDNISOLONE *AE; HEMATEMESIS *AE; MELENA
*AE; GASTROENTEROPATHY *AE; HEMORRHAGE *AE; ANEMIA *AE;
WARFARIN *DI; ASPIRIN *DI; CORTICOSTEROIDS *PT; PREDNISOLONE
*RN; DI *PT; AE *PT

CAS REGISTRY NO.: 50-24-8
[28] THEOPHYLLINE *DI; THEOPHYLLINE *AE; NAUSEA *AE; EMESIS *AE;
GASTROENTEROPATHY *AE; DEHYDRATION *AE; ALLOPURINOL *DI;
RANITIDINE *DI; BRONCHODILATORS *PT; VASODILATORS *PT;
CARDIANTS *PT; DIURETICS *PT; ANTIASTHMATICS *PT;
PHOSPHODIESTERASE-INHIBITORS *PT; THEOPHYLLINE *RN; DI
*PT; AE *PT

[29] *PT
ALLOPURINOL *DI; ALLOPURINOL *AE; NAUSEA *AE; EMESIS *AE;
GASTROENTEROPATHY *AE; THEOPHYLLINE *DI; ANTIULCERS *PT;
ANTIHEURATICS *PT; ALLOPURINOL *RN; DI *PT; AE *PT

CAS REGISTRY NO.: 315-30-0
[30] RANITIDINE *DI; RANITIDINE *AE; ANOREXIA *AE; NAUSEA *AE;
EMESIS *AE; GASTROENTEROPATHY *AE; THEOPHYLLINE *DI;
ANTIULCERS *PT; RANITIDINE *RN; DI *PT; AE *PT

CAS REGISTRY NO.: 66357-35-5
[31] METHOTREXATE *DI; METHOTREXATE *AE; MUCOSITIS *AE;
TRIMETHOPRIM *DI; SULFAMETHOXAZOLE *DI; CYTOSTATICS *PT;
METHOTREXATE *RN; DI *PT; AE *PT

CAS REGISTRY NO.: 59-05-2
[32] GLIBENCLAMIDE *DI; GLIBENCLAMIDE *AE; HYPOGLYCEMIA *AE;
CARBOHYDRATE-METAB.DISORDER *AE; CONFUSION *AE;
MENTAL-DISORDER *AE; METFORMIN *DI; ANTIIDIABETICS *PT;
GLIBENCLAMIDE *RN; DI *PT; AE *PT

CAS REGISTRY NO.: 10238-21-8
[33] METFORMIN *DI; METFORMIN *AE; HYPOGLYCEMIA *AE;
CARBOHYDRATE-METAB.DISORDER *AE; CONFUSION *AE;
MENTAL-DISORDER *AE; GLIBENCLAMIDE *DI; ANTIIDIABETICS *PT;
METFORMIN *RN; DI *PT; AE *PT

CAS REGISTRY NO.: 657-24-9
[34] PROMETHAZINE *DI; PROMETHAZINE *AE; DYSTONIA *AE; MYOPATHY
*AE; EXTRAPYRAMIDAL-DISORDER *AE; ENCEPHALOPATHY *AE;
PROCHLORPERAZINE *DI; ANTIHISTAMINES-H1 *PT; SEDATIVES *PT;
PROMETHAZINE *RN; DI *PT; AE *PT

CAS REGISTRY NO.: 60-87-7
[35] PROCHLORPERAZINE *DI; PROCHLORPERAZINE *AE; DYSTONIA *AE;
MYOPATHY *AE; EXTRAPYRAMIDAL-DISORDER *AE; ENCEPHALOPATHY
*AE; PROMETHAZINE *DI; PSYCHODERATIVES *PT; NEUROLEPTICS *PT;
ANTIEMETICS *PT; DOPAMINE-ANTAGONISTS *PT; PROCHLORPERAZINE *RN; DI
*PT; AE *PT

CAS REGISTRY NO.: 58-38-6
[36] MORPHINE *DI; MORPHINE *AE; CONFUSION
*AE; MENTAL-DISORDER *AE; DROWSINESS *AE; OXYCODONE
*DI; ANALGESICS *PT; NARCOTICS *PT; SEDATIVES *PT;
MORPHINE *RN; DI *PT; AE *PT

CAS REGISTRY NO.: 57-27-2
[37] OXYCODONE *DI; OXYCODONE *AE; CONFUSION
*AE; MENTAL-DISORDER *AE; DROWSINESS *AE; MORPHINE
*DI; ANALGESICS *PT; NARCOTICS *PT; SEDATIVES *PT;
OXYCODONE *RN; DI *PT; AE *PT

CAS REGISTRY NO.: 76-42-6
[38] BROMOCRIPTINE *DI; BROMOCRIPTINE *AE; CONFUSION *AE;
MENTAL-DISORDER *AE; DROWSINESS *AE; THIOPRIMIDINE *DI;
ANTI-PARKINSONISTS *PT; PROLACTIN-ANTAGONISTS *PT;
DOPAMINERGICS *PT; BROMOCRIPTINE *RN; DI *PT; AE *PT

CAS REGISTRY NO.: 25614-03-3
[39] THIOPRIMIDINE *DI; THIOPRIMIDINE *AE; CONFUSION *AE;
MENTAL-DISORDER *AE; DROWSINESS *AE; BROMOCRIPTINE *DI;
PSYCHODERATIVES *PT; NEUROLEPTICS *PT; DOPAMINE-ANTAGONISTS
*PT; CALMODULIN-ANTAGONISTS *PT; THIOPRIMIDINE *RN; DI
*PT; AE *PT

CAS REGISTRY NO.: 50-52-2
[40] FIELD AVAIL.: AB; LA; CT
FILE SEGMENT: Literature

ACCESSION NUMBER: 1988-10343 DRUGU T P S Full-text
TITLE: Pain and Analgesics.
AUTHOR: Kurz H von
LOCATION: Munich, Germany, West
SOURCE: Dtsch.Apoth.Ztg. (127, No. 52-53, 2747-57, 1987) 6 Fig. 10
Tab. 20 Ref.

CODE: DAZE2 ISSN: 0011-9857
AVAIL. OF DOC.: Walther-Streubl-Institut fuer Pharmakologie und Toxikologie
Muessbaumstrasse 26, 8000 Muenchen 2, West Germany.
LANGUAGE: German
DOCUMENT TYPE: Journal

ABSTRACT:

The use of analgesics in the relief of pain is reviewed with reference to the opioids and NSAID, their indications, mechanism of activity, pharmacokinetics, dosage, side effects and interactions with other drugs. Agents that can elicit attacks of asthma, that interact with salicylates and that can be present in analgesic combinations without having analgesic properties are listed. The most serious dangers of using opioids are respiratory paralysis after high doses of addiction following chronic use. NSAID have few side effects when taken sensibly, though they can occasionally induce asthma, Lyell's syndrome and possibly Reye's syndrome.

SECTION HEADING: I Therapeutic
P Pharmacology
S Adverse Effects

CLASSIF. CODE: 8 Pharmacokinetics
35 Adverse Reactions
43 Analgesics, NSAIDs
44 Narcotics
66 Drug Interactions
69 Reviews

CONTROLLED TERM:

[01] PAIN *TR; ANALGESIC *PT; REVIEW *PT; CASES *PT; IN-VIVO *PT
ANALGESICS *PT; MAIN-TOPIC *PT; TR *PT; PH *PT; DM *PT; AE
*PT; DI *PT

[02] PHENACETIN *PH; PHENACETIN *DI; PARACETAMOL *PH; PARACETAMOL *DM; PARACETAMOL *DI; PARACETAMOL *AE; BUCETIN *TR; BUCETIN *DI; BUCETIN *DM; BUCETIN *AE; BUCETIN *PH; PROPYPHENAZONE *TR; PROPYPHENAZONE *AE; ISOPROPRIN *PH; ISOPROPYLPHENAZONE *DM; PROPYPHENAZONE *PH; PROPYPHENAZONE *TR; PHENAZONE *AE; PHENAZONE *DM; PHENAZONE *DI; PHENAZONE *PH; ISOPROPRIN *TR; ISOPROPYLPHENAZONE *TR; ISOPROPRIN *AE; ISOPROPYLPHENAZONE *AE; ISOPROPRIN *DM; ISOPROPYLPHENAZONE *DI; METAMIZOLE *TR; METAMIZOLE *PH; METAMIZOLE *DI; METAMIZOLE *DM; METAMIZOLE *AE; IBUPROFEN *TR; IBUPROFEN *PH; IBUPROFEN *DM; IBUPROFEN *AE; IBUPROFEN *DI; AZAPROAZONE *AE; DICLOFENAC *AE; TR *PT; AE *PT; PH
*PT; DI *PT; DM *PT

[03] SALICYLATE *DM; SALICYLATE *AE; SALICYLATE *DI; ASPIRIN *TR;
ASPIRIN *AE; ASPIRIN *PH; ASPIRIN *DM; ASPIRIN *DI;
SALICYLAMIDE *TR; SALICYLAMIDE *PH; SALICYLAMIDE *AE;
SALICYLAMIDE *DM; SALICYLAMIDE *DI; ETHENZAMIDE *TR;
ETHENZAMIDE *AE; ETHENZAMIDE *PH; ETHENZAMIDE *DI;

	100 mg. corn starch (for mixing) 15 mg. corn starch (for paste) 15 mg. and magnesium stearate 10 mg.
IT	Anabolic agents Analgesics Antiarthritics Antirheumatic agents Arthritis Behcet's syndrome Cholinergic agonists Combination chemotherapy Gout Human Osteoarthritis Pain Rheumatoid arthritis Bercovalis Selective estrogen receptor modulators Tranquilizers (oral combination of strontium salt and other agents for improvement in treatment of arthritic diseases and associated pain)
IT	Drug delivery systems (oral; oral combination of strontium salt and other agents for improvement in treatment of arthritic diseases and associated pain)
IT	Drug delivery systems (tablets; oral combination of strontium salt and other agents for improvement in treatment of arthritic diseases and associated pain)
IT	50-02-2, Dexamethasone 50-18-0, Cyclophosphamide 50-23-7, Hydrocortisone 50-24-8, Prednisolone 50-33-9, Phenylbutazone, biological studies 50-78-2, Aspirin 52-67-5, Penicillamine 53-03-2, Prednisone 53-06-5, Cortisone 53-86-1, Indometacin 56-53-1 57-27-3, Morphine, biological studies 57-42-1, Meperidine 58-15-1, Aminopyrine 59-05-2, Methotrexate 60-80-0, Antipyrine 61-68-7, Mefenamic acid 62-44-2, Phenacetin 62-75-9, Dimethylnitrosoamine 64-85-7, Deoxycortone 67-98-1, Ethamoxtrifelol 69-72-7D, Salicylic acid, derivs. 76-42-6, Oxycodeine 76-57-3, Codeine 76-99-3, Methadone 77-07-6, Levorphanol 77-17-8, Normeperidine 83-43-2, Methylprednisolone 91-64-5, Coumarin, derivs. 118-42-3, Hydroxychloroquine 124-94-7, Trimethacinolone 125-29-1, Hydrocodone 127-31-1, Fludrocortisone 129-20-4, Oxphenbutazone 147-93-3, Thiomalicic acid 359-83-1, Pentazocine 378-44-9, Betamethasone 437-38-7, Fentanyl 446-72-0, Genistein 446-86-6, Azathioprine 466-99-9, Hydromorphone 493-08-3D, Chroman, derivs. 526-26-1, Strontium salicylate 530-78-9, Plufenamic acid 552-94-3, Salaslate 553-39-9, Allenic acid 561-27-3, Heroin 564-25-0, Doxycycline 569-57-3, Chlorotriieniene 644-62-2 853-34-9, Kebuzone 864-19-9, Strontium tartrate 911-45-5, Clomiphene 1400-61-9, Mysterin 1845-11-0, Nafoxidine 2624-43-3, Cyclophenyl 2809-21-3, 3416-24-8, Glucosamine 4419-39-0, Beclomethasone 5104-49-4, Flurbiprofen 5630-53-5, Tibolone 9002-64-6, Parathyroid hormone 9002-72-6, Growth hormone 9004-61-9, Hyaluronic acid 9007-28-7, Chondroitin sulfate 10118-90-8, Minocycline 10448-84-7, Nitromifene 10540-29-1, Tamoxifen 10596-23-3, Clodronate 12244-57-4 13598-36-2D, Phosphonic acid, alkylidenobis, derivs. 15307-86-5, Diclofenac 15687-27-1, Ibuprofen 15722-48-2, Olazalazine 16067-69-9 16088-89-4, 20594-83-6, Nalbuphine 21256-18-8, Omaprozin 22071-15-4, Ketoprofen 22204-53-1, Naproxen 22494-42-4, Diflunisal 25322-68-3D, Polyethylene glycol conjugates with IL-1 receptor derivs. 26171-23-3, Tolmetin 26983-52-8, Diphenol 27303-92-5, Tramadol 27540-07-4 29031-19-4, Glucosamine sulfate 35679-58-1, Fenoprofen 31477-60-8, Ormeloxifene 33369-31-2, Zomepirac 34816-55-2, Moxestrol 36322-90-4,

ER, ES, FI, FR, GB, GR, HU, ID, IT, LT, LU, MC, NL, PL, PT, RO, SE, SI, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, NL, PW, ML, MR, NE, SN, TD, TG	US 200612274 US 20060608 US 2005-269289 20051107
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ED Entered STN: 30 Dec 2005
AB Methods for improving pain management in a mammal, the methods comprising administering a combination of a strontium-containing compound and a second therapeutically and/or prophylactically active substance selected from the group consisting of analgesic agents, anti-inflammatory agents and palliative agents to the mammal. Pharmaceutical compns. for use in such methods, comprising a strontium-containing compound and a second therapeutically and/or prophylactically active substance selected from the group consisting of analgesic agents, anti-inflammatory agents and palliative agents. For example, a tablet containing naproxen 250, strontium malonate 210, lactose 100, corn starch 30, and magnesium stearate 10 mg was formulated.

IT **Antiarthritis**
 Behcet's syndrome
 Bone, neoplasm
 Gout
Osteoarthritis
Rheumatoid arthritis
 Sarcoidosis
Surgery
 (pain associated with; method of improving medical treatment of pain by
 administering combination of strontium-containing compound and second
 active substance)

IT **Drug delivery systems**
 (tablet; method of improving medical treatment of pain by
 administering combination of strontium-containing compound and
 second active substance)

IT 50-33-9, Phenylbutazone, biological studies 50-48-6, Amitriptyline
 50-52-2, Thioridazine 50-53-3, Thorazine, biological studies 50-78-2,
 Aspirin 52-86-8, Haldol 53-86-1, Indometacin 56-06-4,
 2,4-Diamino-6-hydroxypyrimidine 57-27-2, Morphine, biological
 studies 57-42-1, Meperidine 58-33-3, Phenergan 58-38-8,
 Prochlorperazine 58-39-9, Trilafon 60-87-7, Promethazine 61-68-7,
 Mefenamic acid 69-23-8, Fluphenazine 76-42-6, Oxycodeone

56-57-3, Codeine 76-93-3, Methadone 77-07-6, Levorphanol 77-17-8, Normeperidine 79-17-4, Aminoguanidine 84-02-6, Compazine 103-90-2, Paracetamol 113-59-7, Taractan 117-89-5, Trifluoperazine 125-29-1, Hydrocodone 130-61-0, Mellaril 151-16-6, S-(2-Aminothioyl)isoathioure 159-83-1, Pentazocine 364-62-5, Metoclopramide 437-38-7, Fentanyl 440-17-5, Stelazine 466-99-3, Hydromorphone 526-26-1, Strontium sebacylate 548-73-2, Inapsine 561-27-3, Heroin 568-19-9, Strontium tartrate 1977-10-2, Loxapine 2034-23-3, PR 038251 2062-78-4, Orap 2149-70-4 2986-19-8, S-Methylmethionine 2986-20-1, S-Ethylthiothioure 4456-77-3, PR 038470 4673-26-1 5104-49-4, Flurbiprofen 5589-33-0, Serentil 5591-45-7, Navane 5786-21-0, Clozair 6913-17-3, S-Isoopropylthiothioure 7232-21-5, Reglan 7416-34-4, Molindone 13539-58-9, Apazone 15307-86-5, Diclofenac 15622-65-8, Mobar 15687-27-1, Ibuprofen 16067-69-7, Strontium benzenesulfonate 16088-89-4 17035-90-4, NG-Monomethyl-L-arginine 20594-83-6, Nalbuphine 22071-15-4, Ketoprofen 22204-53-1, Naproxen 22780-54-7, 2-Iminopiperidine 22780-54-7D, strontium salts 27203-92-5, Tramadol 27833-64-3, Loxitane 29679-58-1, Fenoprofen 32672-69-8, Mesoridazine besylate 36322-90-4, Piroxicam 37041-91-1, Isoservelle 38194-50-2, Sulindac 40182-75-0, Strontium citrate 40472-00-2 41839-80-9 42408-82-2, Butorphanol 51803-78-2, Nimesulide 52485-79-7, Superoxime 53468-55-8, Dezocine 53774-63-3 58493-49-5, Olvanil 65195-50-8, Scutigeral 71125-38-7, Meloxicam 78754-81-1, PR 191863 8002-04-2, CP55940 106266-05-2, Risperdal 111974-69-7, Quetiapine 111974-72-2, Serquel 123663-49-0, T-614 128007-31-8, Arvanil 132539-06-1, Zyprexa 133587-00-5 133587-00-5, NGMonomethyl-L-arginine acetate 135459-87-9, Strontium ranelate 146939-27-9, Dromen 155836-52-5, BH2731046 156719-41-4, S-Methyl-L-thiocitruilline 158089-95-3, S-2474 159860-31-8, SNC-121 175033-16-0, NCX4016 179469-40-0 181313-72-4, Strontium malonate 183293-82-5 189954-66-3, DPLP 188470-84-7, Parecoxib 188470-85-8, Dynastat 251362-87-5 278172-05-7 303730-87-2 322766-10-9, Tiracoxib 472981-92-3, SB-366791 507471-56-9 535974-91-5 630395-06-1, SVT 2016 769104-84-2 756104-86-4 7596104-90-0 796842-36-9 796842-37-0 796842-38-1 872049-77-9 872049-78-0 872049-79-1 872049-83-7 872049-83-7 872049-85-9 872049-86-0 872049-88-2 872049-89-3 872049-90-6 872049-91-7 872049-92-8 872049-93-9 872049-94-0 872049-95-1 872125-28-5 872200-43-6 872340-66-6, AZD 4717

RL: THU (Therapeutic use); BIOL (Biological study); USGS (Uses)
(method of improving medical treatment of pain by administering combination of strontium-containing compound and second active substance)

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RN: BW, CH, GM, KE, LS, MM, MZ, NA, SD, SL, SZ, TZ, UG, ZM, ZW, AM, AZ, BY, KO, KZ, MD, RU, TJ, TM, AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HU, IE, IS, IT, LT, LU, MC, NL, PL, PT, RO, SR, SI, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GO, GW, MU, MR, NS, SN, TD, TG

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AB The present invention relates to an active substance combination including at least one substituted carbinal compound and at least one non-steroidal anti-inflammatory drug (NSAID), a medicament including the active substance combination, a pharmaceutical formulation including the active substance combination and the use of the active substance combination for the manufacture of a medicament.

TI Pharmaceutical active substance combination comprising substituted carbinal compounds and non-steroidal anti-inflammatory drugs

ST pharmaceutical combination substituted carbinal compd. nonsteroidal antiinflammatory drug

IT Urogenital system
(-related pain; pharmaceutical active substance combination comprising substituted carbinal compds. and non-steroidal anti-inflammatory drugs)

IT Inflammation
(Crohn's disease; pharmaceutical active substance combination comprising substituted carbinal compds. and non-steroidal anti-inflammatory drugs)

IT Intestinal disease
(Crohn's; pharmaceutical active substance combination comprising substituted carbinal compds. and non-steroidal anti-inflammatory drugs)

IT Disease, animal
(Hodgkin's disease; pharmaceutical active substance combination comprising substituted carbinal compds. and non-steroidal anti-inflammatory drugs)

IT Disease, animal
(Peutz-Jegher syndrome; pharmaceutical active substance combination comprising substituted carbinal compds. and non-steroidal anti-inflammatory drugs)

IT Disease, animal
(Sclerodema; pharmaceutical active substance combination comprising substituted carbinal compds. and non-steroidal anti-inflammatory drugs)

IT Drugs of abuse
(abuse of treatment and prevention of; pharmaceutical active substance combination comprising substituted carbinal compds. and non-steroidal anti-inflammatory drugs)

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IT Pain
(acute; pharmaceutical active substance combination comprising substituted carbinal compds. and non-steroidal anti-inflammatory drugs)

IT Carcinoma
(adenocarcinoma; pharmaceutical active substance combination comprising substituted carbinal compds. and non-steroidal anti-inflammatory drugs)

IT Swelling, biological
(after injury; pharmaceutical active substance combination comprising substituted carbinal compds. and non-steroidal anti-inflammatory drugs)

IT Transplant and Transplantation
(allograft, transplant, cornea, rejection; pharmaceutical active substance combination comprising substituted carbinal compds. and non-steroidal anti-inflammatory drugs)

IT Heart, disease
(angina pectoris, pain; pharmaceutical active substance combination comprising substituted carbinal compds. and non-steroidal anti-inflammatory drugs)

IT Blood vessel, neoplasm
(angioblastoma, nasopharynx; pharmaceutical active substance combination comprising substituted carbinal compds. and non-steroidal anti-inflammatory drugs)

IT Inflammation
Spinal column, disease
(ankylosing spondylitis; pharmaceutical active substance combination comprising substituted carbinal compds. and non-steroidal anti-inflammatory drugs)

IT Anemia (disease)
(aplastic; pharmaceutical active substance combination comprising substituted carbinal compds. and non-steroidal anti-inflammatory drugs)

IT Disease, animal
(arthropathy, Bursitis; pharmaceutical active substance combination comprising substituted carbinal compds. and non-steroidal anti-inflammatory drugs)

IT Disease, animal
(back pain, lower, chronic pain; pharmaceutical active substance combination comprising substituted carbinal compds. and non-steroidal anti-inflammatory drugs)

IT Body, anatomical
(back, disease, pain, lower, chronic pain; pharmaceutical active substance combination comprising substituted carbinal compds. and non-steroidal anti-inflammatory drugs)

IT Pain
(back, lower, chronic pain; pharmaceutical active substance combination comprising substituted carbinal compds. and non-steroidal anti-inflammatory drugs)

IT Skin, neoplasm
(basal cell carcinoma; pharmaceutical active substance combination comprising substituted carbinal compds. and non-steroidal anti-inflammatory drugs)

IT Carcinoma
(basal cell; pharmaceutical active substance combination comprising substituted carbinal compds. and non-steroidal anti-inflammatory drugs)

IT Injury
(bone, pain; pharmaceutical active substance combination comprising substituted carbinal compds. and

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non-steroidal anti-inflammatory drugs)

IT Bronchi, disease
(bronchitis; pharmaceutical active substance combination comprising substituted carbinal compds. and non-steroidal anti-inflammatory drugs)

IT Epithelium
(cancer affecting; pharmaceutical active substance combination comprising substituted carbinal compds. and non-steroidal anti-inflammatory drugs)

IT Lip
(cancer; pharmaceutical active substance combination comprising substituted carbinal compds. and non-steroidal anti-inflammatory drugs)

IT Drug delivery systems
(capsules; pharmaceutical active substance combination comprising substituted carbinal compds. and non-steroidal anti-inflammatory drugs)

IT Ischemia
(cardiac; pharmaceutical active substance combination comprising substituted carbinal compds. and non-steroidal anti-inflammatory drugs)

IT Pain
(central, post-operative; pharmaceutical active substance combination comprising substituted carbinal compds. and non-steroidal anti-inflammatory drugs)

IT Uterus, neoplasm
(cervix, carcinoma; pharmaceutical active substance combination comprising substituted carbinal compds. and non-steroidal anti-inflammatory drugs)

IT Carcinoma
(cervix; pharmaceutical active substance combination comprising substituted carbinal compds. and non-steroidal anti-inflammatory drugs)

IT Pain
(chronic; pharmaceutical active substance combination comprising substituted carbinal compds. and non-steroidal anti-inflammatory drugs)

IT Headache
(cluster; pharmaceutical active substance combination comprising substituted carbinal compds. and non-steroidal anti-inflammatory drugs)

IT Intestine, neoplasm
(colon; pharmaceutical active substance combination comprising substituted carbinal compds. and non-steroidal anti-inflammatory drugs)

IT Eye, disease
Inflammation
(conjunctivitis; pharmaceutical active substance combination comprising substituted carbinal compds. and non-steroidal anti-inflammatory drugs)

IT Eye
(cornea, allotransplant, rejection; pharmaceutical active substance combination comprising substituted carbinal compds. and non-steroidal anti-inflammatory drugs)

IT Transplant rejection
(corneal; pharmaceutical active substance combination comprising substituted carbinal compds. and non-steroidal anti-inflammatory drugs)

IT Bladder, disease

IT Inflammation

IT Pain
(dental; pharmaceutical active substance combination comprising substituted carbinal compds. and non-steroidal anti-inflammatory drugs)

IT Mental and behavioral disorders
(depression; pharmaceutical active substance combination comprising substituted carbinal compds. and non-steroidal anti-inflammatory drugs)

IT Eye, disease
(diabetic retinopathy; pharmaceutical active substance combination comprising substituted carbinal compds. and non-steroidal anti-inflammatory drugs)

IT Joint, anatomical
(disease, Bursitis; pharmaceutical active substance combination comprising substituted carbinal compds. and non-steroidal anti-inflammatory drugs)

IT Viscere
(disease, pain; pharmaceutical active substance combination comprising substituted carbinal compds. and non-steroidal anti-inflammatory drugs)

IT Joint, anatomical
(disease, sprain; pharmaceutical active substance combination comprising substituted carbinal compds. and non-steroidal anti-inflammatory drugs)

IT Tendon
(disease, tendinitis; pharmaceutical active substance combination comprising substituted carbinal compds. and non-steroidal anti-inflammatory drugs)

IT Drug delivery systems
(dragees; pharmaceutical active substance combination comprising substituted carbinal compds. and non-steroidal anti-inflammatory drugs)

IT Drug delivery systems
(drops; pharmaceutical active substance combination comprising substituted carbinal compds. and non-steroidal anti-inflammatory drugs)

IT Uterus, disease
(endometriosis; pharmaceutical active substance combination comprising substituted carbinal compds. and non-steroidal anti-inflammatory drugs)

IT Drug delivery systems
(enteric-coated; pharmaceutical active substance combination comprising substituted carbinal compds. and non-steroidal anti-inflammatory drugs)

IT Alcohols, biological studies
RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
(fatty; pharmaceutical active substance combination comprising substituted carbinal compds. and non-steroidal anti-inflammatory drugs)

IT Ulcer
(gastric; pharmaceutical active substance combination comprising substituted carbinal compds. and non-steroidal anti-inflammatory drugs)

IT Inflammation

IT Stomach, disease
(gastritis; pharmaceutical active substance

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combination comprising substituted carbinol compds. and non-steroidal anti-inflammatory drugs)

IT Drug delivery systems (gels; pharmaceutical active substance combination comprising substituted carbinol compds. and non-steroidal anti-inflammatory drugs)

IT Gingiva, disease Inflammation (gingivitis; pharmaceutical active substance combination comprising substituted carbinol compds. and non-steroidal anti-inflammatory drugs)

IT Drug delivery systems (granules; pharmaceutical active substance combination comprising substituted carbinol compds. and non-steroidal anti-inflammatory drugs)

IT Bladder, disease (incontinence; pharmaceutical active substance combination comprising substituted carbinol compds. and non-steroidal anti-inflammatory drugs)

IT Disease, animal (infantile hemangiomas; pharmaceutical active substance combination comprising substituted carbinol compds. and non-steroidal anti-inflammatory drugs)

IT Intestine, disease (inflammatory; pharmaceutical active substance combination comprising substituted carbinol compds. and non-steroidal anti-inflammatory drugs)

IT Drug delivery systems (injections, i.m.; pharmaceutical active substance combination comprising substituted carbinol compds. and non-steroidal anti-inflammatory drugs)

IT Drug delivery systems (injections, i.p.; pharmaceutical active substance combination comprising substituted carbinol compds. and non-steroidal anti-inflammatory drugs)

IT Drug delivery systems (injections, i.v.; pharmaceutical active substance combination comprising substituted carbinol compds. and non-steroidal anti-inflammatory drugs)

IT Drug delivery systems (injections, s.c.; pharmaceutical active substance combination comprising substituted carbinol compds. and non-steroidal anti-inflammatory drugs)

IT Bone, disease (injury, pain; pharmaceutical active substance combination comprising substituted carbinol compds. and non-steroidal anti-inflammatory drugs)

IT Autoimmune disease (insulin-dependent diabetes mellitus; pharmaceutical active substance combination comprising substituted carbinol compds. and non-steroidal anti-inflammatory drugs)

IT Diabetes mellitus (insulin-dependent; pharmaceutical active substance combination comprising substituted carbinol compds. and non-steroidal anti-inflammatory drugs)

IT Drug delivery systems (intratracheal; pharmaceutical active substance combination comprising substituted carbinol compds. and non-steroidal anti-inflammatory drugs)

IT Intestine, disease

(irritable bowel syndrome; pharmaceutical active substance combination comprising substituted carbinol compds. and non-steroidal anti-inflammatory drugs)

IT Heart, disease (ischemia; pharmaceutical active substance combination comprising substituted carbinol compds. and non-steroidal anti-inflammatory drugs)

IT Rheumatoid arthritis (juvenile; pharmaceutical active substance combination comprising substituted carbinol compds. and non-steroidal anti-inflammatory drugs)

IT Drug delivery systems (liquef.; pharmaceutical active substance combination comprising substituted carbinol compds. and non-steroidal anti-inflammatory drugs)

IT Angiogenesis (mediated disorder; pharmaceutical active substance combination comprising substituted carbinol compds. and non-steroidal anti-inflammatory drugs)

IT Neoplasia (metastasis; pharmaceutical active substance combination comprising substituted carbinol compds. and non-steroidal anti-inflammatory drugs)

IT Hydrocarbon waxes, biological studies RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses) (microcryst.; pharmaceutical active substance combination comprising substituted carbinol compds. and non-steroidal anti-inflammatory drugs)

IT Headache (migraine; pharmaceutical active substance combination comprising substituted carbinol compds. and non-steroidal anti-inflammatory drugs)

IT Drug delivery systems (mucosal, transmucosal; pharmaceutical active substance combination comprising substituted carbinol compds. and non-steroidal anti-inflammatory drugs)

IT Drug delivery systems (nasal; pharmaceutical active substance combination comprising substituted carbinol compds. and non-steroidal anti-inflammatory drugs)

IT Pharynx (nasopharynx, angiofibroma; pharmaceutical active substance combination comprising substituted carbinol compds. and non-steroidal anti-inflammatory drugs)

IT Glaucoma (disease) (neovascular; pharmaceutical active substance combination comprising substituted carbinol compds. and non-steroidal anti-inflammatory drugs)

IT Angiogenesis (neovascularization, eye; pharmaceutical active substance combination comprising substituted carbinol compds. and non-steroidal anti-inflammatory drugs)

IT Eye, disease (neovascularization; pharmaceutical active substance combination comprising substituted carbinol compds. and non-steroidal anti-inflammatory drugs)

IT Kidney, disease (nephrotic syndrome; pharmaceutical active substance combination comprising substituted carbinol compds. and non-steroidal anti-inflammatory drugs)

IT Nerve, disease Pain (neuralgia, Herpes; pharmaceutical active substance combination comprising substituted carbinol compds. and non-steroidal anti-inflammatory drugs)

IT Inflammation (neurogenic; pharmaceutical active substance combination comprising substituted carbinol compds. and non-steroidal anti-inflammatory drugs)

IT Nerve, disease (neuropathy, pain; pharmaceutical active substance combination comprising substituted carbinol compds. and non-steroidal anti-inflammatory drugs)

IT Nerve (nociceptive, pain; pharmaceutical active substance combination comprising substituted carbinol compds. and non-steroidal anti-inflammatory drugs)

IT Anti-inflammatory agents (nonsteroidal; pharmaceutical active substance combination comprising substituted carbinol compds. and non-steroidal anti-inflammatory drugs)

IT Drug delivery systems (oral; pharmaceutical active substance combination comprising substituted carbinol compds. and non-steroidal anti-inflammatory drugs)

IT Burn Head and Neck Parturition Sunburn Surgery Tooth, disease (pain; pharmaceutical active substance combination comprising substituted carbinol compds. and non-steroidal anti-inflammatory drugs)

IT Drug delivery systems (parenteral; pharmaceutical active substance combination comprising substituted carbinol compds. and non-steroidal anti-inflammatory drugs)

IT Drug delivery systems (pellets; pharmaceutical active substance combination comprising substituted carbinol compds. and non-steroidal anti-inflammatory drugs)

IT Artery, disease Inflammation (periarteritis nodosa; pharmaceutical active substance combination comprising substituted carbinol compds. and non-steroidal anti-inflammatory drugs)

IT Arm Leg (phantom limb pain; pharmaceutical active substance combination comprising substituted carbinol compds. and non-steroidal anti-inflammatory drugs)

IT Analgesics Anti-infective agents Anti-inflammatory agents Antirheuristics Antiesthmatics Antidepressants Antidiabetic agents Antirheumatic agents

Antitumor agents
Antiucler agents
Antiviral agents
Arthritis
Asthma
Beeswax
Behcet's syndrome
Bladder, neoplasm
Blood vessel, disease
Bone, neoplasm
Brain, neoplasm
Carcinoma
Common cold
Dermatitis
Digestive tract, disease
Digestive tract, neoplasm
Dysmenorrhea
Eczema
Edema
Fever and Hyperthermia
Gelation agents
Gout
Headache
Inflammation
Influenza
Liver, neoplasm
Lung, neoplasm
Mammary gland, neoplasm
Myasthenia gravis
Myositis
Neoplasm
Opioid antagonists
Osteoarthritis
Ovary, neoplasm
Pancreas, neoplasm
Plasticizers
Prostate gland, neoplasm
Psoriasis
Rheumatic fever
Rheumatoid arthritis
Sarcoidosis
Skin, disease
Skin, neoplasm
Strain
(pharmaceutical active substance combination comprising substituted carbinol compds. and non-steroidal anti-inflammatory drugs)

IT Carnauba wax
Fats and Glyceride oils, biological studies
Fatty acids, biological studies
Opioids
Polymers, biological studies
RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses) (pharmaceutical active substance combination comprising substituted carbinol compds. and non-steroidal anti-inflammatory drugs)

IT Myositis (polymyositis; pharmaceutical active substance combination comprising substituted carbinol compds. and non-steroidal anti-inflammatory drugs)

IT Drug delivery systems
(rectal; pharmaceutical active substance combination comprising substituted carbinol compds. and non-steroidal anti-inflammatory drugs)

IT Kidney, neoplasms
(renal cell carcinoma; pharmaceutical active substance combination comprising substituted carbinol compds. and non-steroidal anti-inflammatory drugs)

IT Carcinoma
(renal cell; pharmaceutical active substance combination comprising substituted carbinol compds. and non-steroidal anti-inflammatory drugs)

IT Eye
(retina, neovascularisation; pharmaceutical active substance combination comprising substituted carbinol compds. and non-steroidal anti-inflammatory drugs)

IT Eye, disease
(retrolental fibroplasia; pharmaceutical active substance combination comprising substituted carbinol compds. and non-steroidal anti-inflammatory drugs)

IT Drug delivery systems
(sols.; pharmaceutical active substance combination comprising substituted carbinol compds. and non-steroidal anti-inflammatory drugs)

IT Spinal column, disease
(spondyloarthropathy; pharmaceutical active substance combination comprising substituted carbinol compds. and non-steroidal anti-inflammatory drugs)

IT Disease, animal
(sprain; pharmaceutical active substance combination comprising substituted carbinol compds. and non-steroidal anti-inflammatory drugs)

IT Carcinoma
(squamous cell; pharmaceutical active substance combination comprising substituted carbinol compds. and non-steroidal anti-inflammatory drugs)

IT Drug delivery systems
(suspensions; pharmaceutical active substance combination comprising substituted carbinol compds. and non-steroidal anti-inflammatory drugs)

IT Drug delivery systems
(sustained release; pharmaceutical active substance combination comprising substituted carbinol compds. and non-steroidal anti-inflammatory drugs)

IT Injury
(swelling; pharmaceutical active substance combination comprising substituted carbinol compds. and non-steroidal anti-inflammatory drugs)

IT Arthritis
Synovial membrane, disease
(synovitis; pharmaceutical active substance combination comprising substituted carbinol compds. and non-steroidal anti-inflammatory drugs)

IT Waxes
RU: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
(synthetic; pharmaceutical active substance combination comprising substituted carbinol compds. and non-steroidal anti-inflammatory drugs)

IT Drug delivery systems
(syrups; pharmaceutical active substance combination

comprising substituted carbinol compds. and non-steroidal anti-inflammatory drugs)

IT Lupus erythematosus
(systemic; pharmaceutical active substance combination comprising substituted carbinol compds. and non-steroidal anti-inflammatory drugs)

IT Drug delivery systems
(tablets; immediate release; pharmaceutical active substance combination comprising substituted carbinol compds. and non-steroidal anti-inflammatory drugs)

IT Drug delivery systems
(tablets; pharmaceutical active substance combination comprising substituted carbinol compds. and non-steroidal anti-inflammatory drugs)

IT Inflammation
(tendinitis; pharmaceutical active substance combination comprising substituted carbinol compds. and non-steroidal anti-inflammatory drugs)

IT Inflammation
Thyroid gland, disease
(thyroiditis; pharmaceutical active substance combination comprising substituted carbinol compds. and non-steroidal anti-inflammatory drugs)

IT Drug delivery systems
(transdermal; pharmaceutical active substance combination comprising substituted carbinol compds. and non-steroidal anti-inflammatory drugs)

IT Stomach, disease
(ulcer; pharmaceutical active substance combination comprising substituted carbinol compds. and non-steroidal anti-inflammatory drugs)

IT Inflammation
Intestine, disease
(ulcerative colitis; pharmaceutical active substance combination comprising substituted carbinol compds. and non-steroidal anti-inflammatory drugs)

IT Bone, disease
(vascular necrosis of bone; pharmaceutical active substance combination comprising substituted carbinol compds. and non-steroidal anti-inflammatory drugs)

IT Infection
(viral; pharmaceutical active substance combination comprising substituted carbinol compds. and non-steroidal anti-inflammatory drugs)

IT Disease, animal
(visceral pain; pharmaceutical active substance combination comprising substituted carbinol compds. and non-steroidal anti-inflammatory drugs)

IT Pain
(visceral; pharmaceutical active substance combination comprising substituted carbinol compds. and non-steroidal anti-inflammatory drugs)

IT 561-27-3, Diacetylmorphine
RU: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
(heroin; pharmaceutical active substance combination comprising substituted carbinol compds. and non-steroidal anti-inflammatory drugs)

IT 50-33-9, Phenylbutazone, biological studies 50-78-2, Acetylsalicylic acid 53-66-1, Indometacin 56-81-5, 1,2,3-Propanetriol, biological studies 57-27-2, Morphine, biological studies 57-42-1,

Pethidine 61-68-7, Mefenamic acid 62-67-9, Nalorphine 65-45-2, Salicylamide 67-56-1, Carbinol, biological studies 68-89-3, Metamizol 71-50-1, Acetate, biological studies 76-41-5, Oxymorphone 76-42-6, Oxycodeine 76-58-4, Echymorphine 77-07-6, Levorphanol 92-43-3, Phenidone 103-90-2, Paracetamol 125-28-0, Dihydrocodeine 125-29-1, Hydrocodone 125-58-6, Levomethadone 129-20-4, Oxyphenbutazone 152-02-3, Levosalphan 288-13-1, Pyrazole 288-32-4, Imaizazole, biological studies 302-41-0, Piritramide 357-56-2, Dextromoramide 359-83-1, Pentazocine 437-38-7, Fentanyl 465-65-6, Naloxone 466-99-9, Hydromorphone 469-62-5, Dextropropoxyphene 469-79-4, Ketobemidone 479-92-5, Propyphenazone 530-78-9, Flufenamic acid 644-62-2, Meclomenamic acid 853-34-9, Kebuzone 915-30-0, Diphenoxylate 938-73-6, Ethenamidine 1477-40-3, Levomethadyl 2210-63-1, Mebuprofene 2438-72-4, Bufexamac 4394-00-7, Niflumic acid 5104-49-4, Flurbiprofen 9004-34-6D, Cellulose, ester 9004-57-3, Ethyl cellulose 14521-96-1, Etorphine 15307-86-5, Diclofenac 15687-27-1, Ibuprofen 16590-41-3, Naltrexone 20594-03-6, Nalbuphine 21256-18-8, Oxaprozin 21363-18-8, Vinomil 22071-15-4, Ketoprofen 22204-53-1, Naproxen 22494-42-4, Diflunisal 26171-23-3, Tramadol 29679-58-1, Fenoprofen 30233-64-8, Glycerol monobenenate 30544-47-9, Etofenamate 30748-29-9, Peprazone 31566-31-1, Glycerol monostearate 33005-95-7, Tiaprofenic acid 34552-84-6, Iaxicam 36322-90-4, Piroxicam 36330-56-0, Fenbufen 38194-50-2, Sulindac 41340-25-4, Etodolac 42408-82-2, Butorphanol 42924-53-8, Nabumetone 51803-78-2, Nimesulide 51931-66-9, Tilidine 52485-79-7, Buprenorphine 53164-05-9, Acemetacin 53179-11-6, Loperamide 53648-55-8, Desoxine 53808-88-1, Lonzolac 54340-58-8, Meptazinol 56030-57-9, Sufentanil 59804-37-4, Tenoxicam 66532-85-2, Propacetamol 70374-39-9, Lornoxicam 71125-38-7, Meloxicam 71195-59-9, Alfenant 74103-06-3, Ketorolac 112344-52-2, Plobufen 122154-30-7 122154-31-8 122154-32-9 122154-33-0 122154-35-2 122154-36-3 122154-37-4 122154-38-5 122154-39-6 122154-40-9 122154-41-0 122154-42-1 122154-43-2 122154-44-3 122154-45-4 122154-46-5 122154-47-6 122154-48-7 122154-51-2 122154-52-3 122154-53-4 122154-55-6 122154-56-7 122154-57-8 122154-60-3 122154-63-0 122154-67-0 122154-68-1 122154-70-5 122154-71-6 122154-72-7 122154-73-8 122154-74-9 122154-75-0 122154-76-1 122154-77-2 122154-78-3 122154-79-4 122154-80-7 122154-81-8 122154-82-9 122154-83-0 122154-84-1 122154-85-2 122154-86-3 122154-87-4 122154-88-5 122154-89-6 122154-90-9 122154-91-0 122154-92-1 122154-93-2 122154-94-3 122154-95-4 122154-96-5 122154-97-6 122154-98-7 122154-99-8 122155-00-4 122155-01-5 122155-02-6 122155-03-7 122155-04-8 122155-05-8 122155-06-0 122155-07-1 122155-08-2 122155-09-3 122155-10-4 122155-11-5 122155-12-6 122155-13-7 122155-14-8 122155-15-9 122155-16-0 122155-17-1 122155-18-4 122155-19-5 122155-20-6 122155-21-7 122155-22-0 122155-23-1 122155-24-2 122155-25-3 122155-26-4 122155-27-5 122155-28-6 122155-29-7 122155-30-0 122155-31-1 122155-32-2 122155-33-3 122155-34-4 122155-35-5 122155-36-6 122155-37-7 122155-38-8 122155-39-9 122155-40-2 122155-41-3 122155-42-4 122155-43-5 122155-44-6 122155-45-7 122155-46-8 122155-47-9 122155-48-0 122155-49-1 122155-50-4 122155-51-5 122155-52-6 122155-53-7 122155-54-8 122155-55-9 122155-56-0 122155-57-1 122155-58-2 122155-59-3 122155-60-4 122155-61-7 122155-62-8 122155-63-9 122155-64-0 122155-65-1 122155-67-3 122155-68-4 122155-69-5 122155-70-8 122155-71-9 122155-72-0 122155-73-1 122155-74-2 122155-75-3 122155-76-4 122155-77-5 122155-78-6 122155-79-7 122155-80-0 122155-81-1 122155-82-2 122155-83-3 122155-84-4 122155-85-5 122155-86-6 122155-87-7 122155-88-8 122155-89-9 122155-90-2 122155-91-3 122155-92-4 122155-93-5 122155-94-6 122175-88-6 122175-89-7 122175-90-0 122175-91-1 122175-92-2

RU: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
(pharmaceutical active substance combination comprising substituted carbinol compds. and non-steroidal anti-inflammatory drugs)

IT 122175-93-3 122175-94-4 122175-95-5 122175-96-6 122175-97-7 122175-98-8 122175-99-9 122176-00-5 122176-01-6 131575-03-6 14-Methoxymetopen 132875-61-7, Remifentanil 142155-43-9 148981-63-9 148981-65-1 162011-91-7, Rofecoxib 169590-42-5, Celecoxib 181695-72-7, Valdecoxib 198470-84-7, Parecoxib 202409-33-4, Etoricoxib 247046-52-2 247046-53-3 247046-54-4 247046-55-5 247046-56-6 247046-57-7 247046-58-8 247046-59-9 247046-60-2 247046-61-3 247046-62-4 247046-63-5 247046-64-6 247046-65-7 247046-66-8 247046-67-9 247046-68-0 247046-69-1 247046-70-4 247046-71-5 247046-72-6 251375-82-3 258329-57-6 258329-59-8 258329-61-2 258329-62-3 258329-64-5 258329-65-6 258329-66-7 258329-67-8 258329-68-9 258329-69-0 258329-70-3 258329-71-4 258329-72-5 258329-73-6 258329-74-7 258329-76-9 258329-77-0 258329-78-1 258329-79-2 2586218-44-2 2586218-45-3 2586218-46-4 2586218-47-5 666218-48-6 666218-49-7

RU: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
(pharmaceutical active substance combination comprising substituted carbinol compds. and non-steroidal anti-inflammatory drugs)

L018 ANSWER 16 OF 19 WPIX COPYRIGHT 2006 THE THOMSON CORP on STN
ACCESSION NUMBER: 2005-296105 [30] WPIX
DOC. NO. C01570 [30]
TITLE: Use of a spirocyclic heterocyclic compound for binding an opioid receptor in the treatment of e.g. pain, gastrointestinal dysfunction
DERMINT CLASS: B02; B03
INVENTOR: AJELOO C M; CHU G; DOLLE R E; GU M; LE BOURDONNEC B; LEISTER L K; TUTHILL P A; ZHOU J O; ZHOU Q J; AJELOO C; DOLLE R; LEISTER L; TUTHILL P; ZHOU J
PATENT ASSIGNEE: (ADOL-1) ADOLOR CORP.; (AJELO-1) AJELOO C M; (CHU-1) CHU G; (DOL-1) DOLLE R E; (GU-1) GU M; (LEB-1) LE BOURDONNEC B; (L018-1) LEISTER L K; (TUTH-1) TUTHILL P A; (ZHOU-1) ZHOU Q J
COUNTRY COUNT: 107
PATENT INFO ADRR.:
PATENT NO. KIND DATE WEEK LA PG MAIN IPC
WO 2005033073 A2 20050414 (200530)* EN 573[0]
US 20050159438 A1 20050721 (200548) EN
EP 1675847 A2 20060705 (200644) EN
APPLICATION DETAILS:
PATENT NO. KIND APPLICATION DATE
WO 2005033073 A2 MO 2004-US32479 20041001
US 20050159438 A1 Provisional US 2003-5078649 20031001
EP 1675847 A1 2004-US32479 20041001
EP 1675847 A2 RP 2004-817130 20041001
EP 1675847 A2 WO 2004-US32479 20041001

PILING DETAILS:

PATENT NO	KIND	PATENT NO
EP 1675847	A2	Based on WO 2005033073 A

PRIORITY APPLN. INFO: US 2003-50764P 20031001
US 2004-957554 20041001

TECH

ORGANIC CHEMISTRY - Preferred Compounds: (I') comprise compounds of formula (I').

R1 and R3 = H, alkyl, alkenyl, alkynyl or aryl;

Z = -N(R5)-;

R2 = H, (cyclo)alkyl, alkenyl, alkynyl, alkylcycloalkyl or heterocycloalkyl;

R1+R3, R1+R2 and R2+R3 = 4-6-membered heterocycloalkyl ring;

R = H or alkyl;

Rb = H, alkyl or aryl;

n = 0-3;

A and B' = H, fluorine or alkyl;

A+B' = a double bond between the carbon atoms to which they are attached;

R4 = -Y-W;

Y = a single bond, C(Ra)(Rb), C(Ra)(Rb)C(Ra)(Rb) or C(Ra)(Rb)C(Ra)C(Ra)(Rb);

W = (hetero)aryl;

X = -CH2- -O-, -S-, -SO-, -SO2- or -N(R5)-;

R5 = H, (cyclo)alkyl, -(CH2)-alkenyl, -(CH2)-alkynyl, aryl, -CORb or -SO2Rb;

J=carbon atom to which it is attached = 6-membered aryl or 5- or 6-membered heteroaryl ring.

Provided that:

(a) when R2 is other than -CH(-C(+O)-ORb)(Ra), then R1+R2 and R2+R3 form 4-8-membered heterocycloalkyl ring; when J taken together with the carbon atoms to which it is attached forms phenyl optionally mono- to tri-substituted by 5-14C alkyl, 1-4C alkyl (both optionally substituted by at least one halo or 1-4C alkyl); halo or OH, W is unsubstituted by phenyl; and phenyl optionally mono- to tri-substituted by halo, 1-6C alkyl, 1-6C alkoxy, and phenyl, phenoxyl, 1,3-benzodioxolyl,

2,2-difluoro-1,3-benzodioxolyl, NH2, -N(1-4C alkyl)2 or pyrrolyl; n is 1; R1 and R3 are H; A+B forms a double bond between atoms to which they are attached; Y is a single bond; and X is -O-; then R2 is other than H or methyl; when taken together with the carbon atoms to which it is attached forms a phenyl ring; W is phenyl optionally mono- to tri-substituted by fluoro, OH, 1-6C alkoxy (optionally substituted by at least one fluoro), 2-6 alkenyloxy or -S-1-4C alkyl; is 1; R1 and R3 are H; A+B forms a double bond between atoms to which they are attached; Y is a single bond; and X is -O-; then R2 is other than H or benzyl; and when J forms a 6-membered aryl ring, then it is substituted with other than pyrimidine-2,4-diamine-methyl-5-yl.

In (I'), the spiro carbon and/or the carbon to which -Y-W is attached (preferably the carbon to which -Y-W is attached, or the spiro carbon and the carbon to which -Y-W is attached) is chiral. Preparation: (I) can be prepared by 37 methods as given in the specification e.g. preparation of (Ia) (where X is CH or NJ) involves:

(a) condensing 2'-hydroxyacetophenone derivative of formula (ii) with 1-Boc-4-piperidone in pyrrolidone derivative of formula (iii) at room temperature to obtain N-Boc-spiro(2H-1-benzopyran-2,4'-piperidine)-4(3H)-one derivative of formula (iv);

(b) converting (iv) into an enol triflate derivative of formula (v) using

N-phenylbis(trifluoromethanesulfonamide) of formula (iv) as a triflating agent; and

(c) coupling (v) by Suzuki type coupling with 4-(N,N-diethylaminocarbonyl)phenyl boronic acid (vi) in ethylene glycol dimethyl ether in the presence of tetrakis triphenylphosphine palladium(0) (10 wt.-% on activated carbon), lithium chloride and aqueous solution of sodium carbonate to obtain a substituted spiro(2H-1-benzopyran-2,4'-piperidine) compound of formula (vii), followed by conversion under acidic conditions.

Ru-Ry = not defined.

PHARMACEUTICALS - Preferred Composition: The composition further comprises an antibiotic, antiviral, antifungal, anti-inflammatory and/or anesthetic. Preferred Drugs: The opioid is selected from 73 drug(s), or their diastereomers, salts or complexes as given in the specification e.g. allylprodine, dextromoramide, eptazocine, fentanyl, ketobemidone, loperamide, lofentanil, myrophine, piritramide, tilidine. The agent for the treatment of neuralgia/neuropathic pain is a mild OTC analgesic, a narcotic analgesic, an antiseizure medication or an anti-depressant. The agent for the treatment of depression is a selective serotonin re-uptake inhibitor, a tricyclic compound, a monoamine oxidase inhibitor or an antidepressant compound belonging to the heterocyclic class. The agent for the treatment of incontinence is an anticholinergic agent, an antispasmodic medication, a tricyclic antidepressant, a calcium channel blocker or a beta agonist. An agent for the treatment of Parkinson's disease is selected from deprenyl, amentadine, levodopa or carbidopa. Preferred Method: The prevention or treatment of pain with (I) further involves administering an opioid. The prevention or treatment of urogenital tract disorder with (I) further involves administering an agent for the treatment of incontinence. The prevention or treatment of depression with (I) further involves administering an agent for the treatment of depression. The prevention or treatment of tremors with (I) further involves administering an antiparkinsonian agent. The production or maintenance of an anesthetic state with (I) further involves administering an anesthetic agent selected from an inhaled anesthetic, a hypnotic, an anxiolytic, a neuromuscular blocker or an opioid.

L218 ANSWER 17 OF 19 WPIX COPYRIGHT 2006 THE THOMSON CORP on STN

ACCESSION NUMBER: 2006-030913 [04] WPIX

DOC. NO. CPI: C2006-011203 [04]

TITLE: Use of opioid controlled release oral dosage form for treating chronic obstructive pulmonary disease

DERWENT CLASS: B02

INVENTOR: FLEISCHER W.; LEYENDECKER P.; REIMER K

PATENT ASSIGNEE: (EURO-N) EURCELTIQUE SA

COUNTRY COUNT: 110

PATENT INFO ABBR.:

PATENT NO	KIND DATE	WEEK	LA	PG	MAIN IPC
EP 1604666	A1 20051214 (200604)*	EN	24	[0]	
WO 2005120507	A1 20051222 (200604)	EN			

APPLICATION DETAILS:

PATENT NO	KIND	APPLICATION	DATE
EP 1604666 A1		EP 2004-13468	20040608
WO 2005120507 A1		WO 2005-EP6155	20050608

CONTROLLED TERM:

Medical Descriptors:

*short course therapy
*cancer pain: CO, complication
*cancer pain: DT, drug therapy
chronic pain: CO, complication
chronic pain: DT, drug therapy
neuropathic pain: CO, complication
neuropathic pain: DT, drug therapy

prospective study

medical audit

analgesic activity

receptor blocking

drug safety

drug efficacy

world health organization

lung cancer

head and neck cancer

breast cancer

skin cancer

prostate cancer

kidney cancer

colorectal cancer

fragility fracture

drowsiness: SI, side effect

confusion: SI, side effect

hallucination: SI, side effect

human

male

female

clinical article

controlled study

aged

adult

article

priority journal

Drug Descriptors:

*ketamine: AB, adverse drug reaction

*ketamine: CB, drug combination

*ketamine: DT, drug therapy

*ketamine: IV, intravenous drug administration

*narcotic analgesic agent: AB, adverse drug reaction

*narcotic analgesic agent: CB, drug combination

*narcotic analgesic agent: DT, drug therapy

*narcotic analgesic agent: IV, intravenous drug administration

nonsteroid antiinflammatory agent: AB, adverse drug reaction

nonsteroid antiinflammatory agent: CB, drug combination

nonsteroid antiinflammatory agent: DT, drug therapy

nonsteroid antiinflammatory agent: IV, intravenous drug administration

steroid: AB, adverse drug reaction

steroid: CB, drug combination

steroid: DT, drug therapy

steroid: IV, intravenous drug administration

10/661458

PRIORITY APPLN. INFO: EP 2004-13468 20040608

TECH

PHARMACEUTICALS - Preferred Dosage: The dosage comprises an opioid agonist (e.g. oxycodone, hydrocodone, hydromorphone, morphine, methadone, oxymorphone, fentanyl or sufentanil in the form of free base or salt) or a mixture of opioid agonist and opioid antagonist (e.g. naloxone, nalbuphine or naloxone in the form of free base or salt). Preferably the dosage comprises oxycodone, morphine or a mixture of oxycodone (10 - 150, preferably 10 - 80 mg) and naloxone (1 - 50 mg). Oxycodone and naloxone are present in a ratio up to 25:1 (preferably up to 20:1, especially 2:1 or 1:1). Preferably amount of oxycodone is higher than that of naloxone. The compounds are released from the dosage in a sustained, invariable or independent manner.

L218 ANSWER 18 OF 19 EMBASE COPYRIGHT (c) 2006 Elsevier B.V. All rights reserved on STN

ACCESSION NUMBER: 2005083426 EMBASE Full-text

TITLE: Prospective audit of short-term concurrent ketamine, opioid and anti-inflammatory ('triple-agent') therapy for episodes of acute on chronic pain.

AUTHOR: Good P.; Tullius P.; Jackson K.; Goodchild C.; Ashby M.

CORPORATE SOURCE: Prof. M. Ashby, Centre for Palliative Care, St. Vincent's Hospital, University of Melbourne, PO Box 2900, Fitzroy, Vic. 3065, Australia. ashby@netspace.net.au

SOURCE: Internal Medicine Journal, (2005) Vol. 35, No. 1, pp. 39-44.

Ref: 33

ISSN: 1444-0903 CODEN: IMJNAK

COUNTRY: Australia

DOCUMENT TYPE: Journal; Article

FILE SEGMENT: 006 Internal Medicine

008 Neurology and Neurosurgery

016 Cancer

037 Drug Literature Index

038 Adverse Reactions Titles

LANGUAGE: English

SUMMARY LANGUAGE: English

ENTRY DATE: Entered STN: 3 Mar 2005

Last Updated on STN: 3 Mar 2005

ABSTRACT: Aim: This prospective audit was undertaken in order to document the analgesic response and adverse effects of concurrent short-term ('burst') triple-agent analgesic (ketamine, an opioid and an anti-inflammatory agent - either steroid or non-steroidal) administration, for episodes of acute on chronic pain. The clinical hypothesis in this study is that better pain control may be obtained by simultaneous multiple target receptor blockade. Method: The response of 18 patients is reported. The pain and analgesic requirement data for the 24 h before starting triple-agent therapy were compared with the last 24 h on the triple-agent therapy. Patients were then classified as responders or non-responders. Results: According to stringent clinical criteria, 12 out of the 18 patients were classified as responders. The response rate was highest for somatic pain (7/9) and appeared to decrease with duration of prior uncontrolled pain. Only four out of the 18 patients reported adverse effects and all of these were minor. Conclusions: The results suggest that this 'burst' triple-agent approach is safe and effective in an inpatient palliative care population during episodes of poorly controlled acute on chronic pain, and warrants further investigation to ascertain whether it gives superior results compared to the 'gold-standard' WHO ladder approach.

steroid: IV, intravenous drug administration
 ketorolac: AE, adverse drug reaction
 ketorolac: CB, drug combination
 ketorolac: DT, drug therapy
 ketorolac: IV, intravenous drug administration
 naproxen: AB, adverse drug reaction
 naproxen: CB, drug combination
 naproxen: DT, drug therapy
 naproxen: IV, intravenous drug administration
 dexamethasone: AE, adverse drug reaction
 dexamethasone: CB, drug combination
 dexamethasone: DT, drug therapy
 dexamethasone: IV, intravenous drug administration
 parecoxib: AE, adverse drug reaction
 parecoxib: CB, drug combination
 parecoxib: DT, drug therapy
 parecoxib: IV, intravenous drug administration
 morphine: AB, adverse drug reaction
 morphine: CB, drug combination
 morphine: DO, drug dose
 morphine: DT, drug therapy
 morphine: IV, intravenous drug administration
 hydromorphone: AB, adverse drug reaction
 hydromorphone: CB, drug combination
 hydromorphone: DO, drug dose
 hydromorphone: DT, drug therapy
 hydromorphone: IV, intravenous drug administration
 oxycodone: AB, adverse drug reaction
 oxycodone: CB, drug combination
 oxycodone: DO, drug dose
 oxycodone: DT, drug therapy
 oxycodone: IV, intravenous drug administration
 prednisolone: AB, adverse drug reaction
 prednisolone: CB, drug combination
 prednisolone: DO, drug dose
 prednisolone: DT, drug therapy
 (ketamine) 1867-66-9, 6740-88-1, 81771-21-3; (ketorolac) 74103-06-3; (naproxen) 32204-53-1, 26159-34-2; (dexamethasone) 50-02-2; (parecoxib) 198470-84-7, 198470-85-8; (morphine) 52-26-6, 57-27-2; (hydromorphone) 466-99-9, 71-68-1; (oxycodone) 124-90-3, 76-42-6; (prednisolone) 50-24-8

CAS REGISTRY NO.:
 74103-06-3; (naproxen) 32204-53-1, 26159-34-2;
 (dexamethasone) 50-02-2; (parecoxib) 198470-84-7,
 198470-85-8; (morphine) 52-26-6, 57-27-2; (hydromorphone)
 466-99-9, 71-68-1; (oxycodone) 124-90-3, 76-42-6;
 (prednisolone) 50-24-8

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 2000116057 EMBASE Full-text
 Managing addiction in advanced cancer patients: Why
 AUTHOR: Passik S.D.; Theobald D.S.
 CORPORATE SOURCE: Dr. S.D. Passik, Community Cancer Care Inc., 115 West 19th Street, Indianapolis, IN 46202, United States
 SOURCE: Journal of Pain and Symptom Management, (2000) Vol. 19, No. 3, pp. 229-234.
 Refs: 6
 ISSN: 0885-3924 CODEN: JPSMEU
 PUBLISHER IDENT.: 8 0885-3924(00)00109-3
 COUNTRY: United States
 DOCUMENT TYPE: Journal; Article
 FILE SEGMENT: 016 Cancer
 032 Psychiatry

036 Health Policy, Economics and Management
 037 Drug Literature Index
 040 Drug Dependence, Alcohol Abuse and Alcoholism
 008 Neurology and Neurosurgery

LANGUAGE: English
 SUMMARY LANGUAGE: English
 ENTRY DATE: Entered STN: 13 Apr 2000
 Last Updated on STN: 13 Apr 2000
 ABSTRACT: The management of addiction in patients with advanced cancer can be time-consuming, labor-intensive, and difficult. Some clinicians believe that it is not worth the effort, due in part to a failure to appreciate the deleterious impact of addiction on palliative care efforts and a view of addiction as intractable in any case. Indeed, it is possible that some clinicians perceive addiction not only fatalistically but, because of common misconceptions, believe that managing or attempting to decrease the patient's use of alcohol or illicit substances would be tantamount to depriving a dying patient of a source of pleasure. In this paper, we argue that managing addiction is an essential aspect of palliative care for chemically-dependent and alcoholic patients. The goal of such efforts is not complete abstinence, but exerting enough control over illicit drug and alcohol use to allow palliative care interventions to decrease suffering. To illustrate this view, we describe two patients with chemical-dependency. We highlight the impact of unchecked substance abuse on patients' perpetuation of their own suffering, the complication of symptom management, the diagnosis and treatment of mood/anxiety disorders, and the effect on the patients' family and caregivers. Copyright (C) 2000 U.S. Cancer Pain Relief Committee.

CONTROLED TERM: Medical Descriptors:
 *addiction
 *cancer patient
 palliative therapy
 drug abuse
 anxiety neurosis: ET, etiology
 anxiety neurosis: DT, drug therapy
 anxiety neurosis: CO, complication
 smoking
 advanced cancer
 alcoholism: TH, therapy
 group therapy
 adenocarcinoma
 pleura metastasis: SU, surgery
 pleura effusion: TH, therapy
 drain
 withdrawal syndrome: PC, prevention
 withdrawal syndrome: DT, drug therapy
 cancer pain: DT, drug therapy
 cancer pain: CO, complication
 patient information
 caregiver
 insomnia: DT, drug therapy
 insomnia: CO, complication
 heroin dependence
 lung cancer: ET, radiotherapy
 human
 male
 case report
 adult
 article
 Drug Descriptors:
 alcohol

illicit drug
 lorazepam: DT, drug therapy
 lorazepam: CB, drug combination
 oxycodone: DT, drug therapy
 oxycodone: CB, drug combination
 paracetamol: DT, drug therapy
 paracetamol: CB, drug combination
 trazodone: DT, drug therapy
 trazodone: CB, drug combination
 fentanyl: DT, drug therapy
 fentanyl: CB, drug combination
 fentanyl: TD, transdermal drug administration
 fentanyl: IV, intravenous drug administration
 (alcohol) 64-17-5; (lorazepam) 846-49-1; (oxycodone)
 124-90-3, 76-42-6; (paracetamol) 103-90-2; (trazodone)
 19794-93-5, 25332-39-2; (fentanyl) 437-38-7

FILE 'HOME' ENTERED AT 11:13:55 ON 14 DEC 2006

SEARCH HISTORY

» d his nofile

(FILE 'HOME' ENTERED AT 09:53:23 ON 14 DEC 2006)

FILE 'CAPLUS' ENTERED AT 09:53:50 ON 14 DEC 2006

D SAVED
 ACT ARN458CAAU/A

L1 1 SEA ABB-ON US2003-661458/APPB
 L2 141 SEA ABB-ON PACE G7/AU
 L3 11003 SEA ABB-ON SMITH M7/AU
 L4 1 SEA ABB-ON L2 AND L3
 D SCAN

FILE 'STNGUIDE' ENTERED AT 09:54:39 ON 14 DEC 2006

FILE 'REGISTRY' ENTERED AT 09:55:54 ON 14 DEC 2006

L5 1 SEA ABB-ON MORPHINE/CN
 L6 1 SEA ABB-ON PENTHETYL/CN
 L7 1 SEA ABB-ON SUPERTANIL/CN
 L8 1 SEA ABB-ON ALFENTANIL/CN
 L9 1 SEA ABB-ON OXYMORPHINE/CN
 L10 1 SEA ABB-ON HYDROMORPHONE/CN
 L11 1 SEA ABB-ON OXYCODONE/CN

FILE 'CAPLUS' ENTERED AT 09:56:07 ON 14 DEC 2006

L12 31087 SEA ABB-ON (L5 OR L6 OR L7 OR L8 OR L9 OR L10)
 L13 1073 SEA ABB-ON L11
 D SCAN L4
 L14 12914 SEA ABB-ON OPIOIDS/CT
 L15 1209 SEA ABB-ON L14(L)KAPPA/OBI
 L16 1944 SEA ABB-ON L14(L)MU/OBI
 L17 56591 SEA ABB-ON ACTONIST*/OBI
 L18 368 SEA ABB-ON L15(L)1
 L19 454 SEA ABB-ON L16(L)1
 L20 19117 SEA ABB-ON RESPIRATORY TRACT/OBI
 L21 76 SEA ABB-ON L20(L)CARCINOMA/OBI
 L22 25232 SEA ABB-ON ASTHMA/OBI
 L23 424 SEA ABB-ON BRONCHIECTASIS/OBI OR BRONCHI?/OBI OR DILATATION/OBI
 OR KARTAGENER/OBI
 L24 28786 SEA ABB-ON TUBERCULOSIS/OBI
 L25 4089 SEA ABB-ON BRONCHITIS/OBI
 L26 120 SEA ABB-ON RESPIRATORY SYSTEM, NEOPLASM/CT
 L27 35560 SEA ABB-ON LUNG, NEOPLASM/CT
 L28 4982 SEA ABB-ON CHRONIC OBSTRUCTIVE PULMONARY/OBI OR COPD/OBI
 L29 7726 SEA ABB-ON BRONCHOPNEUMONIA/OBI OR PNEUMONIA/OBI
 L30 136 SEA ABB-ON LARYNgeITIS/OBI
 L31 1101 SEA ABB-ON SINUSITIS/OBI
 L32 2601 SEA ABB-ON EMPHYSEMA/OBI
 L33 6378 SEA ABB-ON FIBROSIS/OBI (L)ALVEOLITIS/OBI OR (PULMONARY/OBI
 OR LUNG/OBI OR RESPIRATORY/OBI) (L) (FIBROSIS/OBI OR SARCOIDOSIS/
 OBI)
 L34 6 SEA ABB-ON SLEEP DISORDERS/CT (L)RESPIRATORY/OBI
 L35 943 SEA ABB-ON SLEEP/OBI (L)APNEA/OBI
 L36 1691 SEA ABB-ON SARCOIDOSIS/CT
 L37 39125 SEA ABB-ON DRUG INTERACTIONS-OLD, NT/CT
 L38 4450 SEA ABB-ON DRUG DELIVERY SYSTEMS-OLD/CT (L)COMB?/OBI

L39 16989 SEA ABB-ON COMBINATION CHEMOTHERAPY/CT
 L40 5480 SEA ABB-ON COMB?/OBI(L) PHARMAC?/OBI
 L41 6 SEA ABB-ON (L12 OR L19) AND (L13 OR L18) AND (L21 OR L22 OR
 L23 OR L24 OR L25 OR L26 OR L27 OR L28 OR L29 OR L30 OR L31 OR
 L32 OR L33 OR L34 OR L35 OR L36) AND (L37 OR L38 OR L39 OR
 L40)
 L42 552 SEA ABB-ON (L12 OR L19) (L)(COMB?/OBI OR COADMIN?/OBI OR
 CODRUG?/OBI OR CONCOMITANT?/OBI OR CONCURRENT?/OBI OR BLEND?/OBI
 I OR MIXTURE?/OBI)
 L43 82 SEA ABB-ON (L13 OR L18) (L)(COMB?/OBI OR COADMIN?/OBI OR
 CODRUG?/OBI OR CONCOMITANT?/OBI OR CONCURRENT?/OBI OR BLEND?/OBI
 I OR MIXTURE?/OBI)
 L44 3 SEA ABB-ON (L12 AND L43) AND (L21 OR L22 OR L23 OR L24 OR L25
 OR L26 OR L27 OR L28 OR L29 OR L30 OR L31 OR L32 OR L33 OR L34
 OR L35 OR L36)
 L45 5 SEA ABB-ON ((L12 AND L43) OR ((L12 OR L19) AND (L13 OR L18)
 AND (L37 OR L38 OR L39 OR L40))) AND (L2 OR L3)

FILE 'EMBASE' ENTERED AT 10:09:18 ON 14 DEC 2006

L46 83 SEA ABB-ON PACE G7/AU
 L47 8120 SEA ABB-ON SMITH M7/AU
 L48 53452 SEA ABB-ON MORPHINE/CT
 E FENTANYL/CT
 E B3-ALL
 L49 26736 SEA ABB-ON FENTANYL/CT OR FENTANYL CITRATE/CT
 E SUFENTANYL/CT
 L50 4395 SEA ABB-ON SUFENTANYL/CT OR SUFENTANYL CITRATE/CT
 E ALFENTANYL/CT
 E B5-ALL
 L51 4482 SEA ABB-ON ALFENTANYL/CT
 E OXYMORPHONE/CT
 L52 805 SEA ABB-ON OXYMORPHONE/CT
 E HYDROMORPHONE/CT
 L53 2957 SEA ABB-ON HYDROMORPHONE/CT
 E OXYCODONE/CT
 L54 3754 SEA ABB-ON OXYCODONE/CT
 E ASTHMA-ALL/CT
 L55 84233 SEA ABB-ON ASTHMA-NT/CT
 E BRONCHIECTASIS-ALL/CT
 L56 4535 SEA ABB-ON BRONCHIECTASIS-NT/CT
 E PULMONARY TUBER/CT
 E E4-ALL
 E E2-ALL
 L57 15140 SEA ABB-ON LUNG TUBERCULOSIS/CT
 E COPD/CT
 E B3-ALL
 E S2-ALL
 L58 26377 SEA ABB-ON CHRONIC OBSTRUCTIVE LUNG DISEASE/CT
 E BRONCHITIS-ALL/CT
 L59 22047 SEA ABB-ON BRONCHITIS-NT/CT
 E BRONCHOPNEUMONIA-ALL/CT
 L60 2275 SEA ABB-ON BRONCHOPNEUMONIA/CT
 E LARYNGETIS-ALL/CT
 L61 2500 SEA ABB-ON LARYNGETIS-NT/CT
 E SINUSITIS-ALL/CT
 L62 12991 SEA ABB-ON SINUSITIS-NT/CT
 E EMPHYSEMA-ALL/CT
 L63 13249 SEA ABB-ON EMPHYSEMA-NT/CT
 E FIBROsing ALV/CT
 E B4-ALL

L64 2738 SEA ABB-ON FIBROsing ALVEOLITIS/CT
 E PULMONARY FIBROSIS/CT
 E S3-ALL
 E S2-ALL
 L65 19537 SEA ABB-ON LUNG FIBROSIS-NT/CT
 E SARCOID/CT
 E SARCOIDOSIS/CT
 E S3-ALL
 E S2-ALL
 L66 11397 SEA ABB-ON SARCOIDOSIS/CT
 E LUNG CANCER/CT
 L67 91685 SEA ABB-ON LUNG CANCER-NT/CT
 E SLEEP APNEA-ALL/CT
 E S2-ALL
 L68 11977 SEA ABB-ON SLEEP APNEA SYNDROME/CT
 L69 20 SEA ABB-ON ((L46 OR L47) OR ((L46 OR L47) AND (L48 OR L49 OR
 L50 OR L51 OR L52 OR L53) AND L54))
 L70 0 SEA ABB-ON ((L46 OR L47) OR ((L47 OR L48 OR L49 OR L50 OR L51 OR L52 OR
 L53) (L)(CB OR IT))
 L71 10397 SEA ABB-ON ((L47 OR L48 OR L49 OR L50 OR L51 OR L52 OR
 L53) (L)(CB OR IT))
 L72 493 SEA ABB-ON L54(L)(CB OR IT)/CT
 L73 5 SEA ABB-ON L71 AND L72 AND (L46 OR L47)
 L74 5 SEA ABB-ON ((L46 AND L47) OR ((L71 AND L72 AND (L46 OR L47))
 D TRIAL 1-5
 L75 38068 SEA ABB-ON DRUG POTENTIATION/CT
 L76 1228 SEA ABB-ON MU OPIATE RECEPTOR AGONIST/CT
 L77 949 SEA ABB-ON KAPPA OPIATE RECEPTOR AGONIST/CT
 L78 208 SEA ABB-ON L76(L)(CB OR IT)/CT
 L79 149 SEA ABB-ON L77(L)(CB OR IT)/CT
 L80 10397 SEA ABB-ON ((L48 OR L49 OR L50 OR L51 OR L52 OR L53) (L)(CB OR
 IT))
 L81 5 SEA ABB-ON ((L46 AND L47) OR ((L80 AND L72 AND (L46 OR L47))
 L82 0 SEA ABB-ON ((L49 OR L49 OR L50 OR L51 OR L52 OR L53 OR L76)
 AND (L77 OR L54)) AND L75 AND (L55 OR L56 OR L57 OR L58 OR L59
 OR L60 OR L61 OR L62 OR L63 OR L64 OR L65 OR L66 OR L67 OR
 L68)
 L83 820 SEA ABB-ON ((L72 OR L79) OR ((L80 OR L78) AND (L55 OR L56 OR
 L57 OR L58 OR L59 OR L60 OR L61 OR L62 OR L63 OR L64 OR L65 OR
 L66 OR L67 OR L68))
 L84 2 SEA ABB-ON ((L72 OR L79) AND ((L80 OR L78) AND (L55 OR L56 OR
 L57 OR L58 OR L59 OR L60 OR L61 OR L62 OR L63 OR L64 OR L65 OR
 L66 OR L67 OR L68))
 D TRIAL 1-2

FILE 'DRUGU' ENTERED AT 10:26:08 ON 14 DEC 2006

L85 2 SEA ABB-ON PACE G7/AU
 L86 1100 SEA ABB-ON SMITH M7/AU
 L87 D TRIAL L85 1-2
 L88 9457 SEA ABB-ON ((L5 OR L6 OR L7 OR L8 OR L9 OR L10)
 E MORPHINE/CT
 L89 19705 SEA ABB-ON MORPHINE/CT
 E FENTANYL/CT
 L90 11240 SEA ABB-ON FENTANYL/CT
 E SUFENTANYL/CT
 L91 2280 SEA ABB-ON SUFENTANYL/CT
 E ALFENTANYL/CT
 L92 2680 SEA ABB-ON ALFENTANYL/CT
 E OXYMORPHONE/CT
 L93 252 SEA ABB-ON OXYMORPHONE/CT
 E HYDROMORPHONE/CT

L114 4 SEA ABB-ON (RABAGO/SDCN OR RACDH7/SDCN OR RAFCO/SDCN OR
 R0654/SDCN OR R16303/SDCN OR 103043-1-0-0/DCSE OR 103043-1-1-0
 /DCSE OR 103043-1-2-0/DCSE OR 758270-1-0-0/DCSE)
 L115 435 SEA ABB-ON L114 OR L113
 L116 513 SEA ABB-ON OXYCODONE/BI,ABEX
 L117 198 SEA ABB-ON MU OPIOID#8/BI,ABEX
 L118 184 SEA ABB-ON KAPPA/BI,ABEX(1W) OPIOID#8/BI,ABEX
 L119 12146 SEA ABB-ON B14-L01/MC OR C14-L01/MC
 L120 100 SEA ABB-ON L117(2A)AGONIST#/BI,ABEX OR ((L117 AND L119)
 L121 102 SEA ABB-ON L118(2A)AGONIST#/BI,ABEX OR ((L118 AND L119)
 L122 466502 SEA ABB-ON ((M782 OR P67?)/M0,M1,M2,M3,M4,M5,M6 OR A61K045/IPC
 OR (B12-C09 OR C12-C09 OR B14-S09 OR C14-S09))
 L123 4 SEA ABB-ON ((L108 OR L109) AND (L111 OR L120)) AND (L115 OR
 L116 OR L121) AND L122
 L124 28604 SEA ABB-ON ASTHMA/BI,ABEX OR BRONCHIECTASIS/BI,ABEX
 OR BRONCHITIS/BI,ABEX(DILATATION/BI,ABEX OR KARTAGENER/BI,ABEX
 OR TUBERCULOSIS/BI,ABEX)
 L125 5007 SEA ABB-ON COPD/BI,ABEX OR CHRONIC OBSTRUCTIVE/BI,ABEX(M) (LUNG
 /BI,ABEX OR PULMONARY/BI,ABEX OR RESPIRATORY/BI,ABEX)
 L126 11360 SEA ABB-ON BRONCHITIS/BI,ABEX OR BRONCHOPNEUMONIA/BI,ABEX
 OR EMPHYSEMA/BI,ABEX
 L127 2462 SEA ABB-ON FIBROsing ALVEOLITIS/BI,ABEX OR FIBROSIS/BI,ABEX
 ((LUNG/BI,ABEX OR PULMONARY/BI,ABEX OR RESPIRATORY/BI,ABEX))
 L128 3624 SEA ABB-ON SARCOIDOSIS/BI,ABEX OR SLEEP APNEA/BI,ABEX
 L129 8806 SEA ABB-ON ((LUNG/BI,ABEX OR PULMONARY/BI,ABEX OR RESPIRATORY/B
 I,ABEX) (2A) (CANCER#/BI,ABEX OR NEOPLAS?/BI,ABEX OR CARCINOMA#/B
 I,ABEX))
 L130 26 SEA ABB-ON ((L111 OR L120) AND (L115 OR L116 OR L121) AND L122
 AND (L124 OR L125 OR L126 OR L127 OR L128 OR L129))
 L131 25 SEA ABB-ON L130 NOT ((L110 OR L123))
 D TRIAL 1-8
 L132 20 SEA ABB-ON SUBANALGES#/BI,ABEX OR SUB ANALGES#/BI,ABEX
 L133 1 SEA ABB-ON L132 AND L130
 D TRIAL
 D KWIC L131 6-10
 D KWIC L131 11-13

FILE 'WPIX' ENTERED AT 10:55:27 ON 14 DEC 2006
 D KWIC L131 11-13

FILE 'WPIX' ENTERED AT 10:56:51 ON 14 DEC 2006
 L134 1 SEA ABB-ON L120 AND L121 AND L122 AND (L124 OR L125 OR L126
 OR L127 OR L128 OR L129)
 L135 20 SEA ABB-ON L115 AND L111 AND L122 AND (L124 OR L125 OR L126
 OR L127 OR L128 OR L129)
 D KWIC L133
 L136 299 SEA ABB-ON ((L111 OR L117)) (5A) ((L116 OR L121)) (5A) ((COMB?/BI,A
 BEX OR CODRUG?/BI,ABEX OR COADMIN?/BI,ABEX OR CONCOMITANT?/BI,A
 BEX OR CONCURRENT?/BI,ABEX OR BLEND?/BI,ABEX OR MIX?/BI,ABEX))
 L137 18 SEA ABB-ON L136 AND L122 AND (L124 OR L125 OR L126 OR L127 OR
 L128 OR L129)
 D QUB
 L138 84 SEA ABB-ON L116(2A)AGONIST#/BI,ABEX
 L139 61 SEA ABB-ON L117(2A)AGONIST#/BI,ABEX
 L140 384 SEA ABB-ON ((L111 OR L139)) (5A) ((L116 OR L138))
 L141 10 SEA ABB-ON L140 (5A) ((COMB?/BI,ABEX OR CODRUG?/BI,ABEX OR
 COADMIN?/BI,ABEX OR CONCOMITANT?/BI,ABEX OR CONCURRENT?/BI,ABEX
 OR BLEND?/BI,ABEX OR MIX?/BI,ABEX))
 L142 2 SEA ABB-ON L141 AND (L124 OR L125 OR L126 OR L127 OR L128 OR

L94 866 SEA ABB-ON HYDROMORPHONE/CT
 E OXYCODONE/CT
 L95 986 SEA ABB-ON OXYCODONE/CT
 E OPIOID AGONIST/CT
 L96 9 SEA ABB-ON ((L85 AND L86) OR ((L85 OR L86) AND (L87 OR L89 OR
 L90 OR L91 OR L92 OR L93 OR L94) AND (L88 OR L95)))
 L97 125676 SEA ABB-ON COMB./CT
 L98 43301 SEA ABB-ON DRUG INTERACTIONS/CC
 L99 86 SEA ABB-ON ((L97 OR L98) AND (L87 OR L89 OR L90 OR L91 OR L92
 OR L93 OR L94) AND (L88 OR L95))
 L100 31287 SEA ABB-ON ASTHMA OR BRONCHIECTASIS OR BRONCHIT? (2A) DILATATION
 OR KARTAGENER OR TUBERCULOSIS
 L101 3808 SEA ABB-ON COPD OR CHRONIC OBSTRUCTIVE(W) (LUNG OR PULMONARY
 OR RESPIRATORY)
 L102 24212 SEA ABB-ON BRONCHITIS OR BRONCHOPNEUMONIA OR PNEUMONIA OR
 LARYNGETIS OR SINUSITIS OR EMPHYSEMA
 L103 1971 SEA ABB-ON FIBROsing ALVEOLITIS OR FIBROSIS(A) (LUNG OR
 PULMONARY OR RESPIRATORY)
 L104 951 SEA ABB-ON SARCOIDOSIS
 L105 17785 SEA ABB-ON ((LUNG OR PULMONARY OR RESPIRATORY) (2A) (CANCER# OR
 NEOPLAS? OR CARCINOMA))
 L106 433 SEA ABB-ON ((L97 OR L98) AND (L87 OR L89 OR L90 OR L91 OR L92
 OR L93 OR L94) AND (L88 OR L95)) AND (L100 OR L101 OR L102 OR
 L103 OR L104 OR L105 OR L106))
 D TRIAL 1-4

FILE 'STNGUIDE' ENTERED AT 10:34:45 ON 14 DEC 2006

FILE 'WPIX' ENTERED AT 10:36:41 ON 14 DEC 2006
 L108 79 SEA ABB-ON PACE G7/AU
 L109 2413 SEA ABB-ON SMITH M7/AU
 L110 1 SEA ABB-ON L108 AND L109
 D TRIAL

FILE 'STNGUIDE' ENTERED AT 10:37:33 ON 14 DEC 2006

FILE 'WPIX' ENTERED AT 10:38:56 ON 14 DEC 2006
 E B04-A04+ALL/MC
 E B07-H+ALL/MC
 E B12-M10A+ALL/MC
 E B12-M10C+ALL/MC
 E B14-C01+ALL/MC
 E B14-H01+ALL/MC
 E B14-H01N+ALL/MC
 E B14-J02+ALL/MC
 E B14-K01+ALL/MC
 E B14-L01+ALL/MC
 E B14-S09+ALL/MC
 FILE 'STNGUIDE' ENTERED AT 10:39:14 ON 14 DEC 2006
 FILE 'WPIX' ENTERED AT 10:41:48 ON 14 DEC 2006
 L111 3147 SEA ABB-ON MORPHINE/BI,ABEX OR FENTANYL/BI,ABEX OR ALFENTANYL/
 BI,ABEX OR SUFENTANYL/BI,ABEX OR OXYMORPHONE/BI,ABEX OR
 MRZ2593/BI,ABEX OR MRZ 2593/BI,ABEX OR HYDROMORPHONE/BI,ABEX
 E OXYCODONE/CN
 L112 4 SEA ABB-ON OXYCODONE?/CN
 L113 431 SEA ABB-ON L112/DCR
 SEL L112 SDRB,SDCN,DCSE

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L129)
 L143 25 SEA ABB=ON L140 AND L122 AND (L124 OR L125 OR L126 OR L127 OR
 L128 OR L129)

FILE 'MEDLINE' ENTERED AT 11:04:27 ON 14 DEC 2006

D SAVED
 ACT ARN458MEDA/A

 L144(94) SEA FILE=+MEDLINE ABB=ON PACE GT/AU
 L145(10732) SEA FILE=+MEDLINE ABB=ON SMITH MT/AU
 L146(0) SEA FILE=+MEDLINE ABB=ON L144 AND L145
 L147(26104) SEA FILE=+MEDLINE ABB=ON MORPHINE/CT
 L148(10382) SEA FILE=+MEDLINE ABB=ON FENTANYL-NT/CT
 L149(294) SEA FILE=+MEDLINE ABB=ON OXYMORPHONE/CT
 L150(704) SEA FILE=+MEDLINE ABB=ON HYDROMORPHONE/CT
 L151(540) SEA FILE=+MEDLINE ABB=ON OXYCODONE/CT
 L152(124991) SEA FILE=+MEDLINE ABB=ON LUNG DISEASES, OBSTRUCTIVE-NT/CT
 L153(5936) SEA FILE=+MEDLINE ABB=ON BRONCHIECTASIS+NT/CT
 L154(57086) SEA FILE=+MEDLINE ABB=ON TUBERCULOSIS, PULMONARY-NT/CT
 L155(3460) SEA FILE=+MEDLINE ABB=ON BRONCHOPNEUMONIA/CT
 L156(3610) SEA FILE=+MEDLINE ABB=ON LARYNGITIS+NT/CT
 L157(11628) SEA FILE=+MEDLINE ABB=ON SINUSITIS-NT/CT
 L158(11372) SEA FILE=+MEDLINE ABB=ON PULMONARY FIBROSIS/CT
 L159(1561) SEA FILE=+MEDLINE ABB=ON SARCOIDOSIS, PULMONARY/CT
 L160(11384) SEA FILE=+MEDLINE ABB=ON LUNG NEOPLASMS+NT/CT
 L161(12706) SEA FILE=+MEDLINE ABB=ON SLEEP APNEA SYNDROMES+NT/CT
 L162(0) SEA FILE=+MEDLINE ABB=ON (L144 OR L145) AND (L147 OR L148 OR L149)
 L163 0 SEA ABB=ON L146 OR L162

 ACT ARN458MED1/A

 L164(28104) SEA FILE=+MEDLINE ABB=ON MORPHINE/CT
 L165(10382) SEA FILE=+MEDLINE ABB=ON FENTANYL-NT/CT
 L166(294) SEA FILE=+MEDLINE ABB=ON OXYMORPHONE/CT
 L167(704) SEA FILE=+MEDLINE ABB=ON HYDROMORPHONE/CT
 L168(540) SEA FILE=+MEDLINE ABB=ON OXYCODONE/CT
 L169(124991) SEA FILE=+MEDLINE ABB=ON LUNG DISEASES, OBSTRUCTIVE-NT/CT
 L170(5936) SEA FILE=+MEDLINE ABB=ON BRONCHIECTASIS+NT/CT
 L171(57086) SEA FILE=+MEDLINE ABB=ON TUBERCULOSIS, PULMONARY-NT/CT
 L172(3460) SEA FILE=+MEDLINE ABB=ON BRONCHOPNEUMONIA/CT
 L173(3610) SEA FILE=+MEDLINE ABB=ON LARYNGITIS+NT/CT
 L174(11628) SEA FILE=+MEDLINE ABB=ON SINUSITIS-NT/CT
 L175(11372) SEA FILE=+MEDLINE ABB=ON PULMONARY FIBROSIS/CT
 L176(1561) SEA FILE=+MEDLINE ABB=ON SARCOIDOSIS, PULMONARY/CT
 L177(11384) SEA FILE=+MEDLINE ABB=ON LUNG NEOPLASMS+NT/CT
 L178(12706) SEA FILE=+MEDLINE ABB=ON SLEEP APNEA SYNDROMES+NT/CT
 L179 1 SEA ABB=ON (L164 OR L165 OR L166 OR L167) AND L168 AND (L169
 OR L170 OR L171 OR L172 OR L173 OR L174 OR L175 OR L176 OR
 L177 OR L178)

 ACT ARN458MED2/A

 L180(28104) SEA FILE=+MEDLINE ABB=ON MORPHINE/CT
 L181(10382) SEA FILE=+MEDLINE ABB=ON FENTANYL-NT/CT
 L182(294) SEA FILE=+MEDLINE ABB=ON OXYMORPHONE/CT
 L183(704) SEA FILE=+MEDLINE ABB=ON HYDROMORPHONE/CT
 L184(540) SEA FILE=+MEDLINE ABB=ON OXYCODONE/CT
 L185(108974) SEA FILE=+MEDLINE ABB=ON DRUG INTERACTIONS+NT/CT
 L186(42787) SEA FILE=+MEDLINE ABB=ON DRUG COMBINATIONS/CT
 L187(97253) SEA FILE=+MEDLINE ABB=ON DRUG THERAPY, COMBINATION/CT

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L188(108974) SEA FILE=+MEDLINE ABB=ON DRUG INTERACTIONS+NT/CT
 L189(42787) SEA FILE=+MEDLINE ABB=ON DRUG COMBINATIONS/CT
 L190(97253) SEA FILE=+MEDLINE ABB=ON DRUG THERAPY, COMBINATION/CT
 L191(1136) SEA FILE=+MEDLINE ABB=ON RECEPTORS, OPIOID, MU/CT(L)AG/CT
 L192(881) SEA FILE=+MEDLINE ABB=ON RECEPTORS, OPIOID, KAPPA/CT(L)AG/CT
 L193(23) SEA FILE=+MEDLINE ABB=ON L193 AND L194 AND (L190 OR L191 OR L192)
 L194(240557) SEA FILE=+MEDLINE ABB=ON DOSE-RESPONSE RELATIONSHIP, DRUG/CT
 L195(1) SEA ABB=ON L195 AND L196 AND CONDITIONING, OPERANT/CT
 L196(1) SEA ABB=ON L195 AND L205 AND L206

 ACT ARN458MED3/A

FILE 'STNGUIDE' ENTERED AT 11:05:43 ON 14 DEC 2006

FILE 'CAPLUS' ENTERED AT 11:06:43 ON 14 DEC 2006

D QUE L1
 D QUE L45
 L208 5 SEA ABB=ON (L1 OR L45)

FILE 'EMBASE' ENTERED AT 11:06:46 ON 14 DEC 2006

D QUE L81

FILE 'DRUGU' ENTERED AT 11:06:47 ON 14 DEC 2006

D QUE L96

FILE 'WPIX' ENTERED AT 11:06:48 ON 14 DEC 2006

D QUE L110

D QUE L123

FILE 'MEDLINE' ENTERED AT 11:06:50 ON 14 DEC 2006

D QUE L163

FILE 'DRUGU, CAPLUS, EMBASE' ENTERED AT 11:07:12 ON 14 DEC 2006

L209 15 DUP REM L96 L208 L81 (4 DUPLICATES REMOVED)

ANSWERS '1-9' FROM FILE DRUGU

ANSWERS '10-13' FROM FILE CAPLUS

ANSWERS '14-15' FROM FILE EMBASE

D IALL ED ABS 1-15

FILE 'STNGUIDE' ENTERED AT 11:07:44 ON 14 DEC 2006

FILE 'CAPLUS' ENTERED AT 11:10:12 ON 14 DEC 2006

D QUE L1

D QUE L45

D IALL 1-10

D IBB ED ABS HIT 11-15

D IBB ABEQ TECH HITSTR 16-17

D IALL 18-19

FILE 'HOME' ENTERED AT 11:13:55 ON 14 DEC 2006

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L210 5 SEA ABB=ON (L1 OR L45)

FILE 'EMBASE' ENTERED AT 11:10:14 ON 14 DEC 2006

D QUE L81

FILE 'DRUGU' ENTERED AT 11:10:15 ON 14 DEC 2006

D QUE L96

FILE 'WPIX' ENTERED AT 11:10:16 ON 14 DEC 2006

D QUE L110

D QUE L123

L211 4 SEA ABB=ON (L110 OR L123)

FILE 'MEDLINE' ENTERED AT 11:10:19 ON 14 DEC 2006

D QUE L163

FILE 'DRUGU, CAPLUS, WPIX, EMBASE' ENTERED AT 11:10:37 ON 14 DEC 2006

L212 16 DUP REM L96 L210 L211 L81 (7 DUPLICATES REMOVED)

ANSWERS '1-9' FROM FILE DRUGU

ANSWERS '10-13' FROM FILE CAPLUS

ANSWER '14' FROM FILE WPIX

ANSWERS '15-16' FROM FILE EMBASE

D IBB ED ABS 1-16

FILE 'STNGUIDE' ENTERED AT 11:11:03 ON 14 DEC 2006

FILE 'CAPLUS' ENTERED AT 11:12:31 ON 14 DEC 2006

D QUE L41

D QUE L44

L213 5 SEA ABB=ON (L41 OR L44) NOT L210

FILE 'EMBASE' ENTERED AT 11:12:33 ON 14 DEC 2006

D QUE L82

D QUE L84

L214 2 SEA ABB=ON L84 NOT L81

FILE 'DRUGU' ENTERED AT 11:12:35 ON 14 DEC 2006

D QUE L107

L215 4 SEA ABB=ON L107 NOT L96

FILE 'WPIX' ENTERED AT 11:12:38 ON 14 DEC 2006

D QUE L134

D QUE L142

L216 2 SEA ABB=ON (L134 OR L142) NOT L211

FILE 'MEDLINE' ENTERED AT 11:12:43 ON 14 DEC 2006

D QUE L189

D QUE L197

D QUE L207

D QUE L179

L217 6 SEA ABB=ON (L189 OR L197 OR L207 OR L179)

FILE 'MEDLINE, DRUGU, CAPLUS, WPIX, EMBASE' ENTERED AT 11:13:15 ON 14 DEC 2006

L218 19 DUP REM L217 L215 L213 L216 L214 (0 DUPLICATES REMOVED)

ANSWERS '1-6' FROM FILE MEDLINE

ANSWERS '7-10' FROM FILE DRUGU

ANSWERS '11-15' FROM FILE CAPLUS

ANSWERS '16-17' FROM FILE WPIX

ANSWERS '18-19' FROM FILE EMBASE